

Welcome to STN International! Enter x:x

LOGINID:ssspta1202txn

PASSWORD :

TERMINAL (ENTER 1, 2, 3, OR ?) :2

* * * * * * * * * * * * * Welcome to STN International * * * * * * *, * * * *

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America
NEWS 2 "Ask CAS" for self-help around the clock
NEWS 3 Feb 24 PCTGEN now available on STN
NEWS 4 Feb 24 TEMA now available on STN
NEWS 5 Feb 26 NTIS now allows simultaneous left and right truncation
NEWS 6 Feb 26 PCTFULL now contains images
NEWS 7 Mar 04 SDI PACKAGE for monthly delivery of multifile SDI results
NEWS 8 Mar 24 PATDPAFULL now available on STN
NEWS 9 Mar 24 Additional information for trade-named substances without structures available in REGISTRY
NEWS 10 Apr 11 Display formats in DGENE enhanced
NEWS 11 Apr 14 MEDLINE Reload
NEWS 12 Apr 17 Polymer searching in REGISTRY enhanced
NEWS 13 Jun 13 Indexing from 1947 to 1956 added to records in CA/CAPLUS
NEWS 14 Apr 21 New current-awareness alert (SDI) frequency in WPIDS/WPINDEX/WPIX
NEWS 15 Apr 28 RDISCLOSURE now available on STN
NEWS 16 May 05 Pharmacokinetic information and systematic chemical names added to PHAR
NEWS 17 May 15 MEDLINE file segment of TOXCENTER reloaded
NEWS 18 May 15 Supporter information for ENCOMPPAT and ENCOMPLIT updated
NEWS 19 May 19 Simultaneous left and right truncation added to WSCA
NEWS 20 May 19 RAPRA enhanced with new search field, simultaneous left and right truncation
NEWS 21 Jun 06 Simultaneous left and right truncation added to CBNB
NEWS 22 Jun 06 PASCAL enhanced with additional data
NEWS 23 Jun 20 2003 edition of the FSTA Thesaurus is now available
NEWS 24 Jun 25 HSDB has been reloaded
NEWS 25 Jul 16 Data from 1960-1976 added to RDISCLOSURE
NEWS 26 Jul 21 Identification of STN records implemented
NEWS 27 Jul 21 Polymer class term count added to REGISTRY
NEWS 28 Jul 22 INPADOC: Basic index (/BI) enhanced; Simultaneous Left and Right Truncation available

NEWS EXPRESS April 4 CURRENT WINDOWS VERSION IS V6.01a, CURRENT MACINTOSH VERSION IS V6.0b(ENG) AND V6.0Jb(JP), AND CURRENT DISCOVER FILE IS DATED 01 APRIL 2003
NEWS HOURS STN Operating Hours Plus Help Desk Availability
NEWS INTER General Internet Information
NEWS LOGIN Welcome Banner and News Items
NEWS PHONE Direct Dial and Telecommunication Network Access to STN
NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may

09/ 830,227

result in loss of user privileges and other penalties.

FILE 'HOME' ENTERED AT 13:11:32 ON 04 AUG 2003

=> file reg
COST IN U.S. DOLLARS . SINCE FILE TOTAL
ENTRY SESSION
FULL ESTIMATED COST . 0.21 0.21

FILE 'REGISTRY' ENTERED AT 13:11:41 ON 04 AUG 2003
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES : 3 AUG 2003 HIGHEST RN 560059-45-2
DICTIONARY FILE UPDATES : 3 AUG 2003 HIGHEST RN 560059-45-2

TSCA INFORMATION NOW CURRENT THROUGH JANUARY 6, 2003

Please note that search-term pricing does apply when conducting SmartSELECT searches.

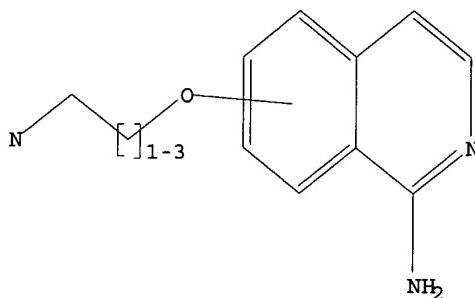
Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details:
<http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf>

=>
Uploading 09830227b.str

L1 STRUCTURE UPLOADED

```
=> d 11  
L1 HAS NO ANSWERS  
L1                      STR
```



Structure attributes must be viewed using STN Express query preparation.

```
=> s 11 ful
FULL SEARCH INITIATED 13:12:02 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED -      8920 TO ITERATE
```

09/ 830,227

100.0% PROCESSED 8920 ITERATIONS
SEARCH TIME: 00.00.01

284 ANSWERS

L2 284 SEA SSS FUL L1

| | | |
|----------------------|------------|---------|
| => file caplus | SINCE FILE | TOTAL |
| COST IN U.S. DOLLARS | ENTRY | SESSION |
| FULL ESTIMATED COST | 148.15 | 148.36 |

FILE 'CAPLUS' ENTERED AT 13:12:06 ON 04 AUG 2003
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 4 Aug 2003 VOL 139 ISS 6
FILE LAST UPDATED: 3 Aug 2003 (20030803/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 12
L3 10 L2

=> d 13 1- ibib abs hitstr
YOU HAVE REQUESTED DATA FROM 10 ANSWERS - CONTINUE? Y/(N):y

L3 ANSWER 1 OF 10 CAPLUS COPYRIGHT 2003 ACS on STN
ACCESSION NUMBER: 2003:376636 CAPLUS
DOCUMENT NUMBER: 138:385436
TITLE: Preparation of 4-(1,1-dioxido-2-
isothiazolidinyl)benzenamines as inhibitors of
blood-coagulation factor Xa for the treatment of
thromboembolic diseases
INVENTOR(S): Dorsch, Dieter; Cezanne, Bertram; Tsaklakidis,
Christos; Mederski, Werner; Gleitz, Johannes; Barnes,
Christopher
PATENT ASSIGNEE(S): Merck Patent GmbH, Germany
SOURCE: PCT Int. Appl., 81 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: German
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2003039543 | A1 | 20030515 | WO 2002-EP11349 | 20021010 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, | | | | |

GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
 LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
 PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
 UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
 TJ, TM

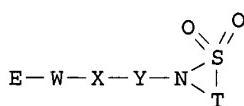
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
 CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
 PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
 NE, SN, TD, TG

DE 10155075 A1 20030522 DE 2001-10155075 20011109

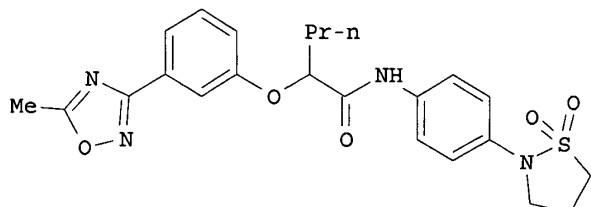
PRIORITY APPLN. INFO.: DE 2001-10155075 A 20011109

OTHER SOURCE(S): MARPAT 138:385436

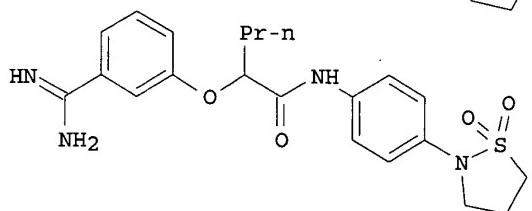
GI



I



II



III

AB Title compds. I [E = (un)substituted aryl, heteroaryl; W = C(R₂)₂, [C(R₂)₂], OC(R₂)₂, etc.; R₂ = H, A, [C(R₃)₂]_n, etc.; R₃ = H, A; X = CONR₂, CONR₂C(R₃)₂, C(R₃)₂NR₂, etc.; Y = alkylene, cycloalkylene, Ar-diyl (sic), etc.; Ar = (un)substituted Ph, naphthyl, biphenyl; T = (un)substituted (CH₂)_p, e.g., N, O, S; n = 0-2; p = 1-6] and their pharmaceutically acceptable salts were prep'd. For example, Raney-Nickel mediated redn. of oxadiazol II, e.g., prep'd. from 4-nitroaniline in 4-steps, afforded isothiazolidine III acetate. In blood-coagulation factor Xa inhibition studies, isothiazolidine III acetate exhibited an IC₅₀ value of 3.5 x 10⁻⁷ M. Compds. I are claimed useful for the treatment of thromboembolic diseases and tumors.

IT 524957-14-0P 524957-15-1P 524957-16-2P

524957-36-6P 524957-37-7P

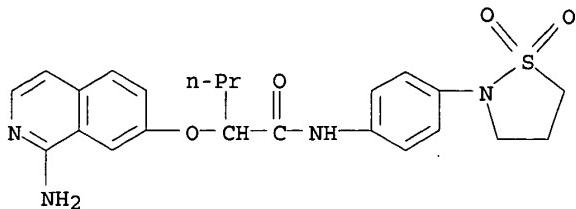
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; prep'n. of isothiazolidinylbenzenamines as inhibitors of blood coagulation factor Xa for the treatment of thromboembolic diseases)

RN 524957-14-0 CAPLUS

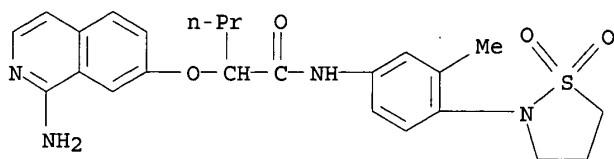
CN Pentanamide, 2-[(1-amino-7-isooquinolinyl)oxy]-N-[4-(1,1-dioxido-2-

isothiazolidinyl)phenyl]- (9CI) (CA INDEX NAME)



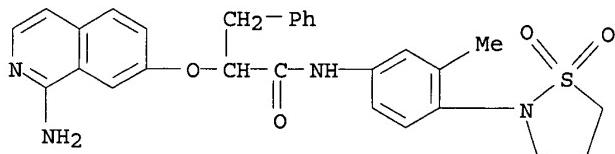
RN 524957-15-1 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isooquinolinyl)oxy]-N-[4-(1,1-dioxido-2-isothiazolidinyl)-3-methylphenyl]- (9CI) (CA INDEX NAME)



RN 524957-16-2 CAPLUS

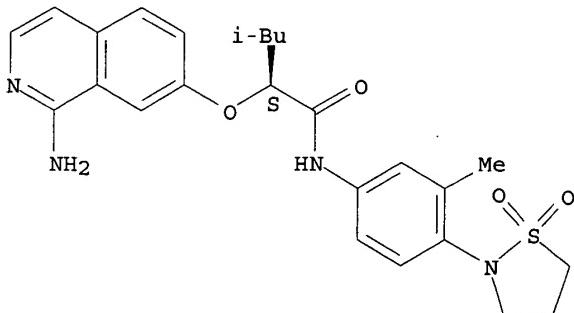
CN Benzenepropanamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[4-(1,1-dioxido-2-isothiazolidinyl)-3-methylphenyl]- (9CI) (CA INDEX NAME)



RN 524957-36-6 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isooquinolinyl)oxy]-N-[4-(1,1-dioxido-2-isothiazolidinyl)-3-methylphenyl]-4-methyl-, (2S)- (9CI) (CA INDEX NAME)

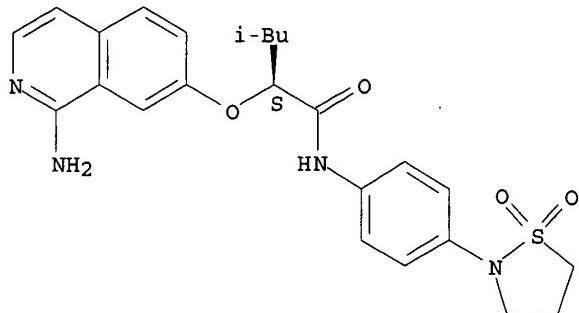
Absolute stereochemistry.



RN 524957-37-7 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isooquinolinyl)oxy]-N-[4-(1,1-dioxido-2-isothiazolidinyl)phenyl]-4-methyl-, (2S)- (9CI) (CA INDEX NAME)

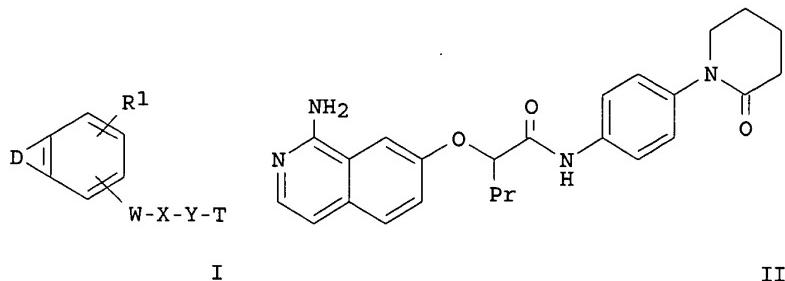
Absolute stereochemistry.



REFERENCE COUNT: 14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 2 OF 10 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 2003:133044 CAPLUS
 DOCUMENT NUMBER: 138:187647
 TITLE: Preparation of phenyl derivatives as coagulation factor Xa inhibitors
 INVENTOR(S): Dorsch, Dieter; Cezanne, Bertram; Tsaklakidis, Christos; Mederski, Werner; Gleitz, Johannes; Barnes, Christopher
 PATENT ASSIGNEE(S): Merck Patent GmbH, Germany
 SOURCE: PCT Int. Appl., 78 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: German
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--|----------|--------------------|----------|
| WO 2003013531 | A1 | 20030220 | WO 2002-EP7798 | 20020712 |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | |
| DE 10139060 | A1 | 20030220 | DE 2001-10139060 | 20010808 |
| PRIORITY APPLN. INFO.: | | | DE 2001-10139060 A | 20010808 |
| OTHER SOURCE(S): | CASREACT 138:187647; MARPAT 138:187647 | | | |
| GI | | | | |



AB Novel Ph compds. I [D = (un)satd. 3 - 4 alkylene chain, contg. 1 - 2 N, O and/or S {may be substituted with halogen, A, {C(R3)2}n-Ar, {C(R3)2}n-Het1, {C(R3)2}n-cycloalkyl, OR2, N(R2)2, NO2, CN, CO2R2, CON(R2)2, NR2COA, NR2SO2A, COR2, SO2NR2, S(O)mA}; W = C(R2)2, {C(R2)2}2, OC(R2)2, NR2C(R2)2; X = CONR2, CONR2C(R3)2, C(R3)2NR2, C(R3)2NR2C(R3)2; Y = alkylene, cycloalkylene, Het-diyl, Ar-diyl; T = (un)substituted heterocycle contg. 1 - 4 of N, O and/or S; A = (un)branched C1-6-alkyl {may contain O, S, CH:CH or substituted with 1 - 7 F}; R1 = H, halogen, A, OR2, N(R2)2, NO2, CN, CO2R2, CON(R2)2, {C(R3)2}nAr, {C(R3)2}n-Het, {C(R3)2}n-cycloalkyl; R2 = H, A, {C(R3)2}nAr, {C(R3)2}n-Het, {C(R3)2}n-cycloalkyl; R3 = H, A; Ar = (un)substituted Ph, naphthyl, biphenyl {may be substituted with halogen, A, OR3, N(R3)2, NO2, CN, CO2R3, CON(R3)2, NR3COA, NR3CON(R3)2, NR3SO2A, COR3, SO2N(R3)2, SOMA}; Het = (un)satd. or arom. heterocycle (contg. 1 - 4 N, O and/or S and may be substituted with halogen, A, {C(R3)2}n-Het1, {C(R3)2}n-cycloalkyl, OR2, N(R2)2, NO2, CN, CO2R2, CON(R2)2, NR2COA, NR2CON(R2)2, NR2SO2A, COR2, SO2NR2, S(O)mA); Het1 = (un)satd. or arom. heterocycle (contg. 1 - 2 N, O and/or S and may be substituted with halogen, A, OR2, N(R2)2, NO2, CN, CO2R2, CON(R2)2, NR2COA, NR2CON(R2)2, NR2SO2A, COR2, SO2NR2, S(O)mA); halogen = Cl Br, F, I; n = 0 - 2; m = 0 - 2] are claimed. I and their pharmaceutically acceptable derivs., solvates, stereoisomers and their mixts., are inhibitors of coagulation factor Xa and can be used in the prophylaxis and/or therapy of thromboembolic diseases and in the treatment of tumors. Thus isoquinoline II was prepnd. from 7-hydroxyisoquinoline via O-alkylation with Me(CH₂)₂CHBrCO₂Et, sapon., amidation with 1-(4-aminophenyl)piperidin-2-one, isoquinoline N-oxidn., isoquinoline N-oxide amination with pyridine, and reaction with ethanolamine. II was tested for thrombin receptor binding ability [IC₅₀ = 3.5 x 10⁻⁷ M vs. FXa; IC₅₀ = 2.2 x 10⁻⁷ M vs. TF]. I was used in the prepn. of drug formulations (injections, suppositories, solns., solvates, tablets, etc.).

| | | | |
|----|--------------|--------------|--------------|
| IT | 498540-34-4P | 498540-36-6P | 498540-56-0P |
| | 498540-57-1P | 498540-59-3P | 498540-60-6P |
| | 498540-61-7P | 498540-62-8P | 498540-63-9P |
| | 498540-64-0P | 498540-65-1P | 498540-66-2P |
| | 498540-67-3P | 498540-68-4P | 498540-69-5P |
| | 498540-70-8P | 498540-72-0P | 498540-73-1P |
| | 498540-74-2P | 498540-75-3P | 498540-76-4P |
| | 498540-77-5P | 498540-78-6P | 498540-79-7P |
| | 498540-80-0P | 498540-81-1P | 498540-82-2P |
| | 498540-83-3P | 498540-84-4P | 498540-85-5P |
| | 498540-86-6P | 498540-87-7P | 498540-88-8P |
| | 498540-89-9P | 498540-90-2P | 498540-91-3P |
| | 498540-92-4P | 498540-93-5P | 498540-94-6P |
| | 498540-95-7P | 498540-96-8P | 498540-97-9P |
| | 498540-98-0P | 498540-99-1P | 498541-00-7P |
| | 498541-01-8P | 498541-02-9P | 498541-03-0P |
| | 498541-04-1P | 498541-05-2P | 498541-06-3P |
| | 498541-07-4P | 498541-08-5P | 498541-29-0P |

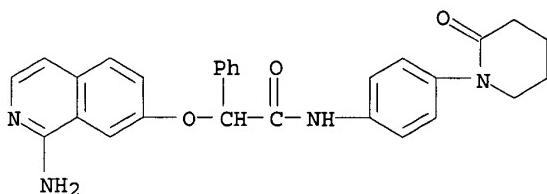
498541-31-4P 498541-33-6P 498541-35-8P
 498541-37-0P 498541-38-1P 498541-39-2P
 498541-56-3P 498541-58-5P 498541-60-9P
 498541-62-1P 498541-64-3P 498541-66-5P
 498541-67-6P 498541-68-7P 498541-69-8P
 498541-70-1P 498541-71-2P 498541-72-3P
 498541-73-4P 498541-74-5P 498541-75-6P
 498541-76-7P 498541-78-9P 498541-80-3P
 498541-82-5P 498541-84-7P 498541-87-0P
 498541-88-1P 498541-89-2P 498541-90-5P
 498541-92-7P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of bicyclic benzene derivs. as coagulation factor Xa inhibitors)

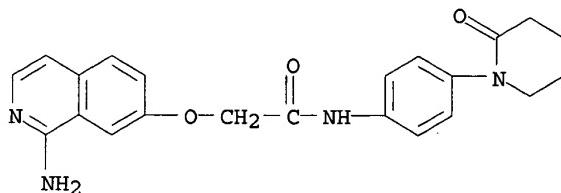
RN 498540-34-4 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498540-36-6 CAPLUS

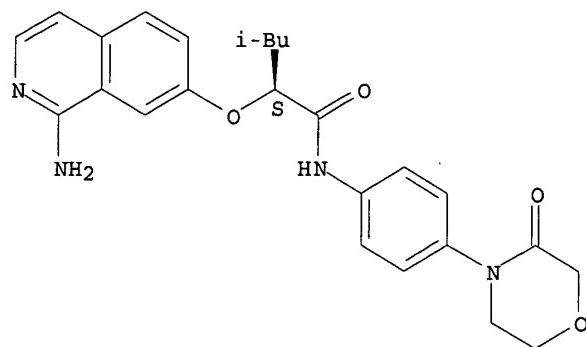
CN Acetamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498540-56-0 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(3-oxo-4-morpholinyl)phenyl]-, (2S)- (9CI) (CA INDEX NAME)

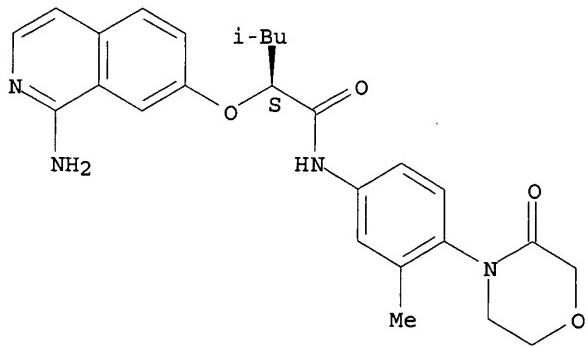
Absolute stereochemistry.



RN 498540-57-1 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[3-methyl-4-(3-oxo-4-morpholinyl)phenyl]-, (2S)- (9CI) (CA INDEX NAME)

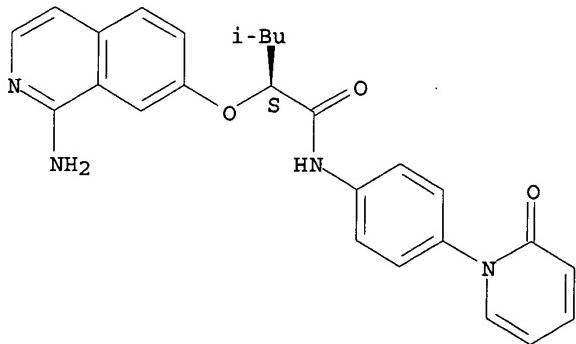
Absolute stereochemistry.



RN 498540-59-3 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

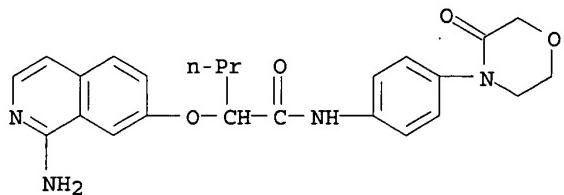


RN 498540-60-6 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(3-oxo-4-

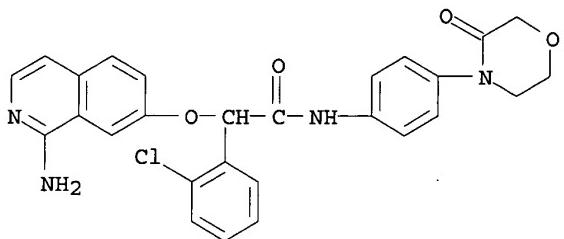
09/ 830,227

morpholinyl)phenyl] - (9CI) (CA INDEX NAME)



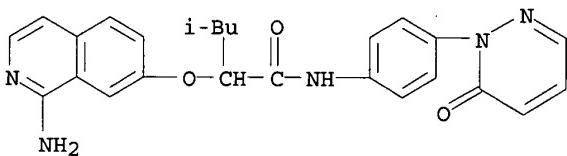
RN 498540-61-7 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyloxy)-2-chloro-N-[4-(3-oxo-4-morpholinyl)phenyl] - (9CI) (CA INDEX NAME)



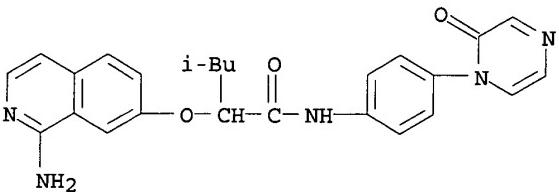
RN 498540-62-8 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyloxy)-4-methyl-N-[4-(6-oxo-1(6H)-pyridazinyl)phenyl] - (9CI) (CA INDEX NAME)



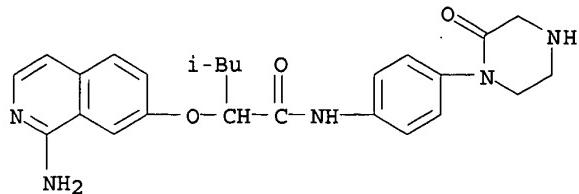
RN 498540-63-9 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyloxy)-4-methyl-N-[4-(2-oxo-1(2H)-pyrazinyl)phenyl] - (9CI) (CA INDEX NAME)

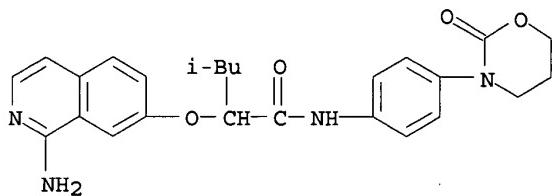


RN 498540-64-0 CAPLUS

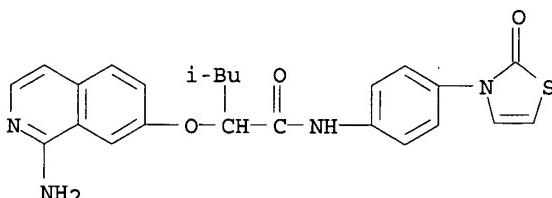
CN Pentanamide, 2-[(1-amino-7-isoquinolinyloxy)-4-methyl-N-[4-(2-oxo-1-piperazinyl)phenyl] - (9CI) (CA INDEX NAME)



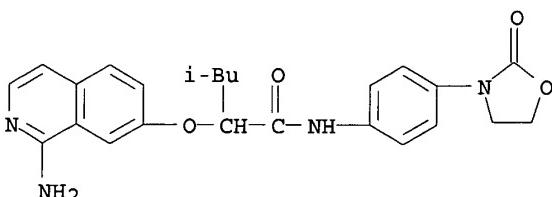
RN 498540-65-1 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(dihydro-2-oxo-2H-1,3-oxazin-3(4H)-yl)phenyl]-4-methyl- (9CI) (CA INDEX NAME)



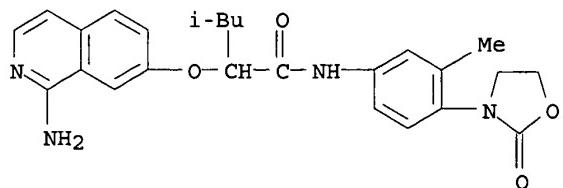
RN 498540-66-2 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(2-oxo-3(2H)-thiazolyl)phenyl]- (9CI) (CA INDEX NAME)



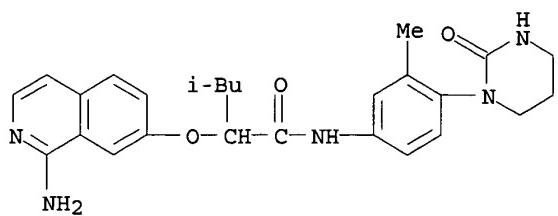
RN 498540-67-3 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(2-oxo-3-oxazolidinyl)phenyl]- (9CI) (CA INDEX NAME)



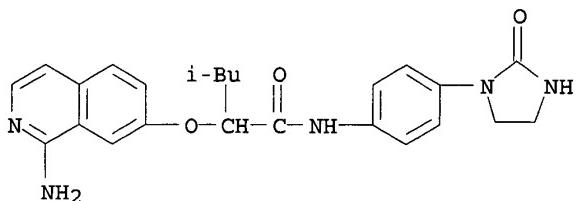
RN 498540-68-4 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[3-methyl-4-(2-oxo-3-oxazolidinyl)phenyl]- (9CI) (CA INDEX NAME)



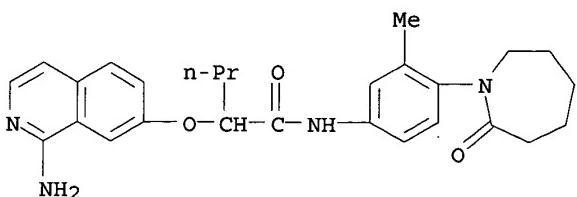
RN 498540-69-5 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[3-methyl-4-(tetrahydro-2-oxo-1(2H)-pyrimidinyl)phenyl]- (9CI) (CA INDEX NAME)



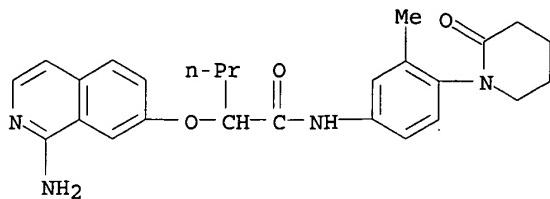
RN 498540-70-8 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(2-oxo-1-imidazolidinyl)phenyl]- (9CI) (CA INDEX NAME)



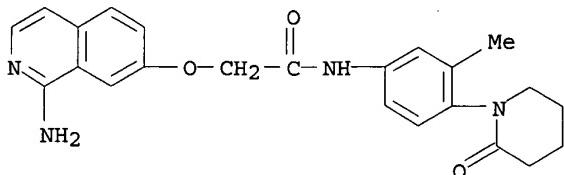
RN 498540-72-0 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(hexahydro-2-oxo-1H-azepin-1-yl)-3-methylphenyl]- (9CI) (CA INDEX NAME)



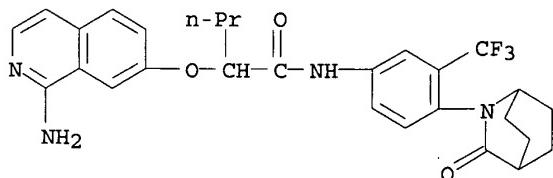
RN 498540-73-1 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



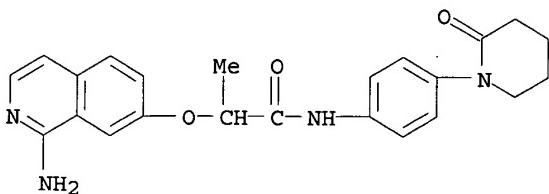
RN 498540-74-2 CAPLUS
CN Acetamide, 2-[(1-amino-7-isquinolinyl)oxy]-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



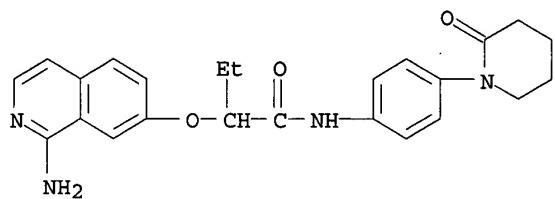
RN 498540-75-3 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isquinolinyl)oxy]-N-[4-(3-oxo-2-azabicyclo[2.2.2]oct-2-yl)-3-(trifluoromethyl)phenyl]- (9CI) (CA INDEX NAME)



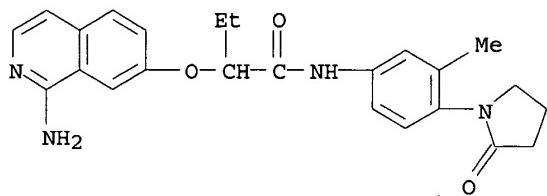
RN 498540-76-4 CAPLUS
CN Propanamide, 2-[(1-amino-7-isquinolinyl)oxy]-N-[4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



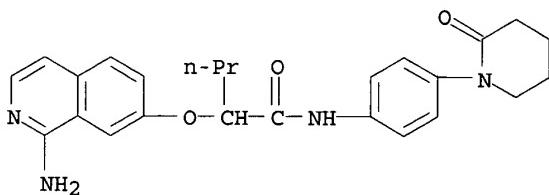
RN 498540-77-5 CAPLUS
CN Butanamide, 2-[(1-amino-7-isquinolinyl)oxy]-N-[4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



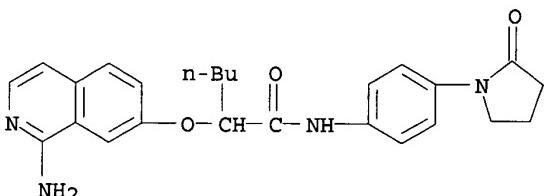
RN 498540-78-6 CAPLUS
CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(2-oxo-1-pyrrolidinyl)phenyl]- (9CI) (CA INDEX NAME)



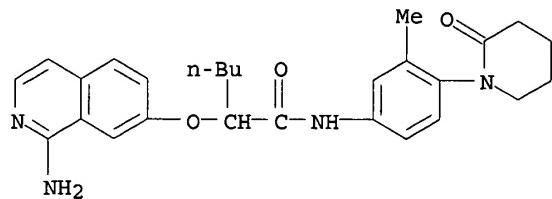
RN 498540-79-7 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



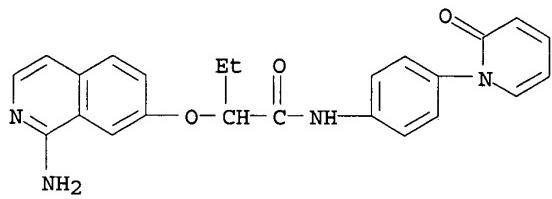
RN 498540-80-0 CAPLUS
CN Hexanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1-pyrrolidinyl)phenyl]- (9CI) (CA INDEX NAME)



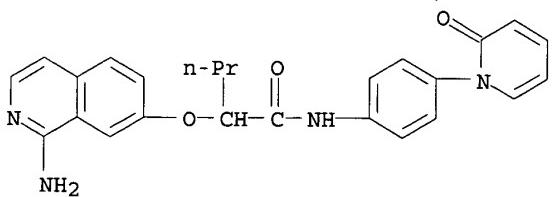
RN 498540-81-1 CAPLUS
CN Hexanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



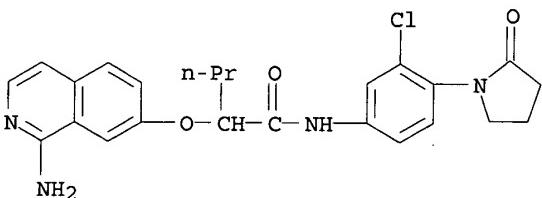
RN 498540-82-2 CAPLUS
CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]- (9CI) (CA INDEX NAME)



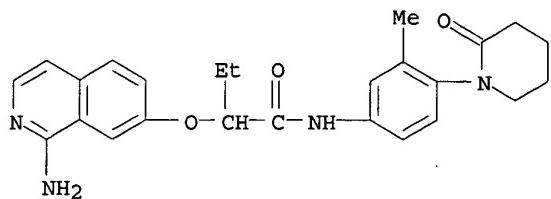
RN 498540-83-3 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498540-84-4 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-chloro-4-(2-oxo-1-pyrrolidinyl)phenyl]- (9CI) (CA INDEX NAME)

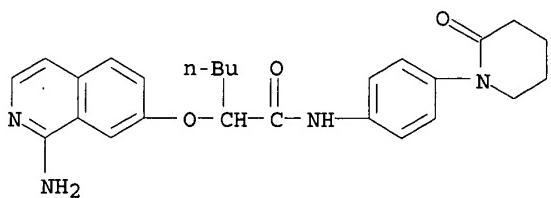


RN 498540-85-5 CAPLUS
CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



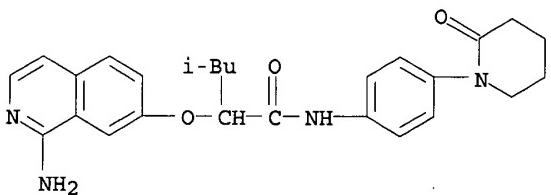
RN 498540-86-6 CAPLUS

CN Hexanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



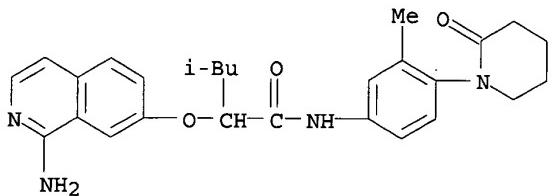
RN 498540-87-7 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498540-88-8 CAPLUS

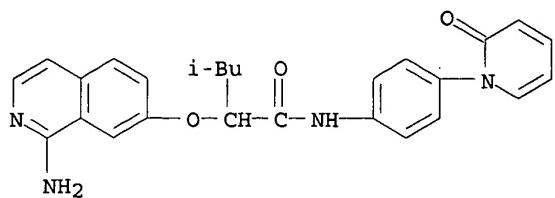
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

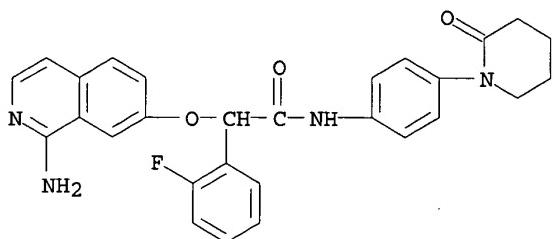
RN 498540-89-9 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]- (9CI) (CA INDEX NAME)



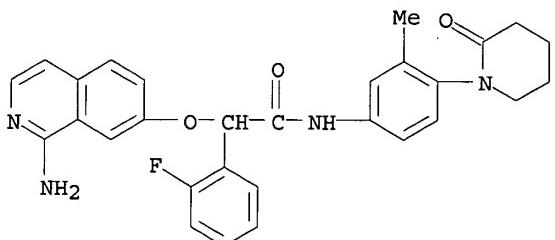
RN 498540-90-2 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-2-fluoro-N-[4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498540-91-3 CAPLUS

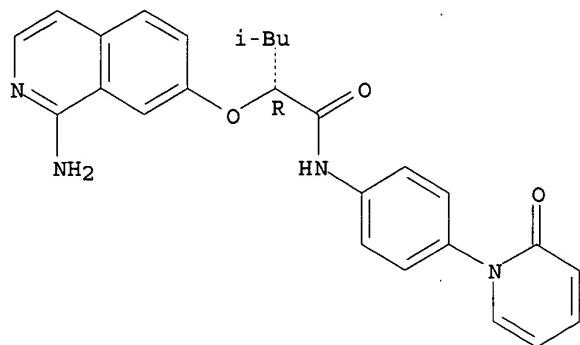
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-2-fluoro-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498540-92-4 CAPLUS

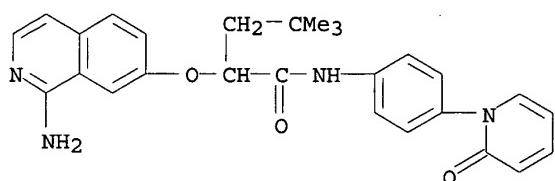
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



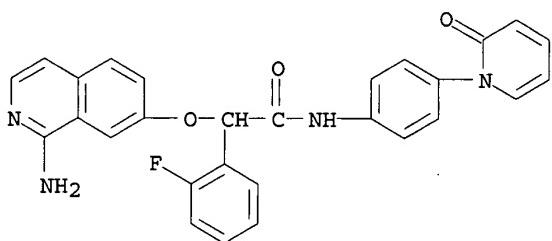
RN 498540-93-5 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyloxy)-4,4-dimethyl-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]- (9CI) (CA INDEX NAME)



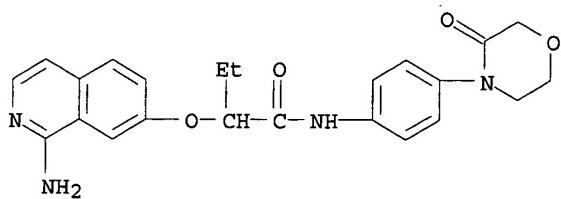
RN 498540-94-6 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyloxy)-2-fluoro-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498540-95-7 CAPLUS

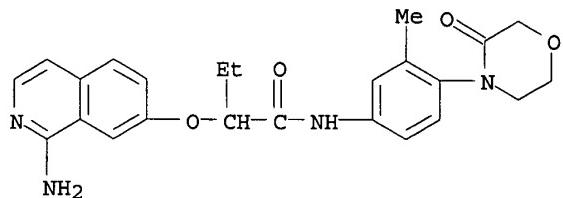
CN Butanamide, 2-[(1-amino-7-isoquinolinyloxy)-N-[4-(3-oxo-4-morpholinyl)phenyl]- (9CI) (CA INDEX NAME)



09/ 830,227

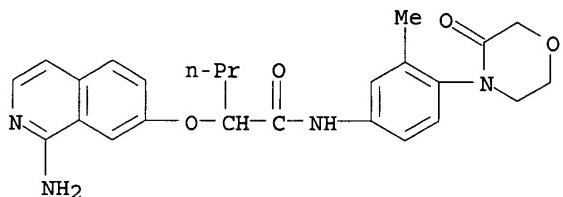
RN 498540-96-8 CAPLUS

CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(3-oxo-4-morpholinyl)phenyl]- (9CI) (CA INDEX NAME)



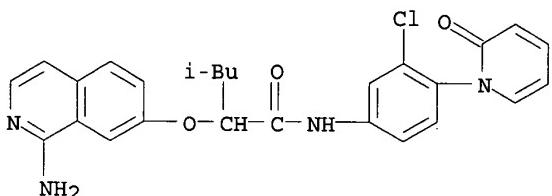
RN 498540-97-9 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(3-oxo-4-morpholinyl)phenyl]- (9CI) (CA INDEX NAME)



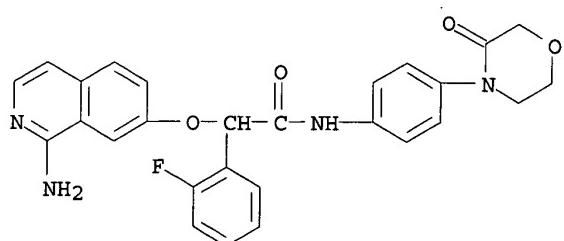
RN 498540-98-0 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-chloro-4-(2-oxo-1(2H)-pyridinyl)phenyl]-4-methyl- (9CI) (CA INDEX NAME)



RN 498540-99-1 CAPLUS

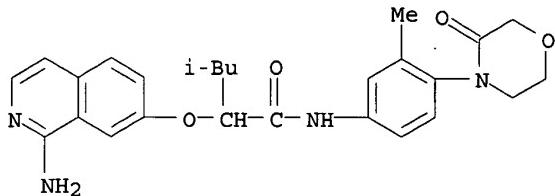
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-2-fluoro-N-[4-(3-oxo-4-morpholinyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498541-00-7 CAPLUS

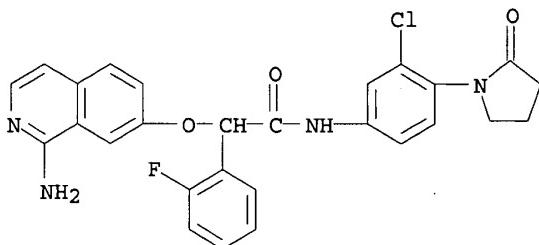
09/ 830,227

CN Pentanamide, 2-[(1-amino-7-isooquinolinyl)oxy]-4-methyl-N-[3-methyl-4-(3-oxo-4-morpholinyl)phenyl]- (9CI) (CA INDEX NAME)



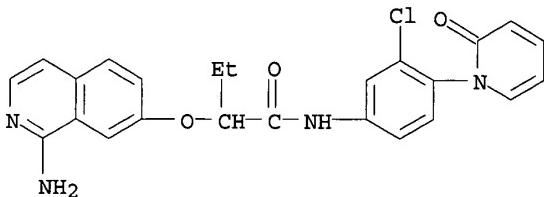
RN 498541-01-8 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[3-chloro-4-(2-oxo-1-pyrrolidinyl)phenyl]-2-fluoro- (9CI) (CA INDEX NAME)



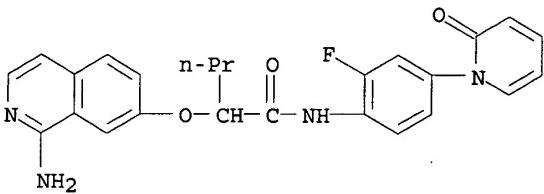
RN 498541-02-9 CAPLUS

CN Butanamide, 2-[(1-amino-7-isooquinolinyl)oxy]-N-[3-chloro-4-(2-oxo-1(2H)-pyridinyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498541-03-0 CAPLUS

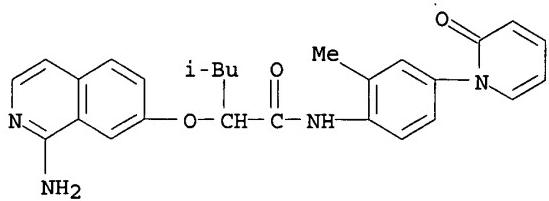
CN Pentanamide, 2-[(1-amino-7-isooquinolinyl)oxy]-N-[2-fluoro-4-(2-oxo-1(2H)-pyridinyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498541-04-1 CAPLUS

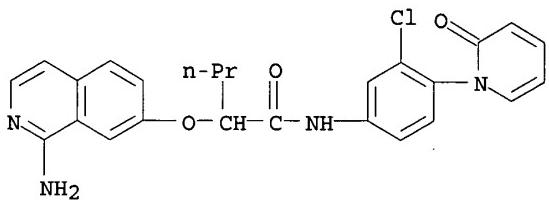
CN Pentanamide, 2-[(1-amino-7-isooquinolinyl)oxy]-4-methyl-N-[2-methyl-4-(2-

oxo-1(2H)-pyridinyl)phenyl]- (9CI) (CA INDEX NAME)



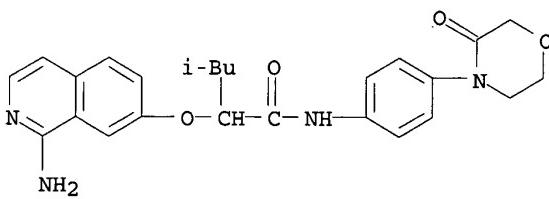
RN 498541-05-2 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-chloro-4-(2-oxo-1(2H)-pyridinyl)phenyl]- (9CI) (CA INDEX NAME)



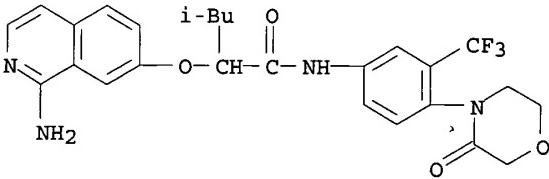
RN 498541-06-3 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(3-oxo-4-morpholinyl)phenyl]- (9CI) (CA INDEX NAME)



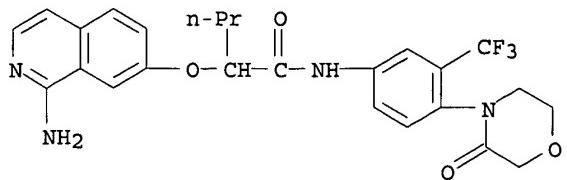
RN 498541-07-4 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(3-oxo-4-morpholinyl)-3-(trifluoromethyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498541-08-5 CAPLUS

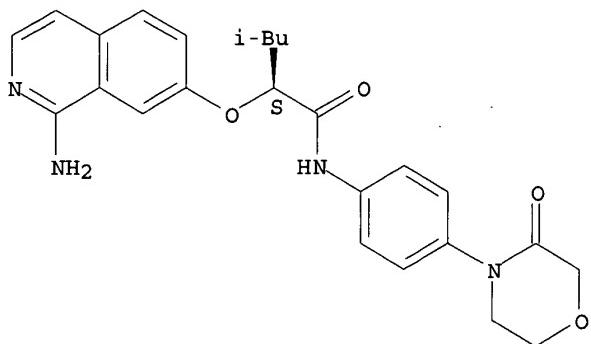
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(3-oxo-4-morpholinyl)-3-(trifluoromethyl)phenyl]- (9CI) (CA INDEX NAME)



RN 498541-29-0 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(3-oxo-4-morpholinyl)phenyl]-, monohydrochloride, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

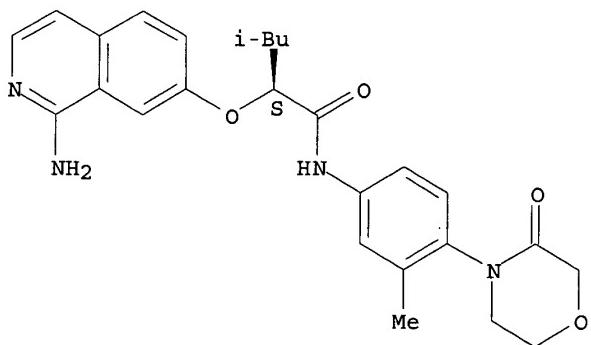


● HCl

RN 498541-31-4 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[3-methyl-4-(3-oxo-4-morpholinyl)phenyl]-, monohydrochloride, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



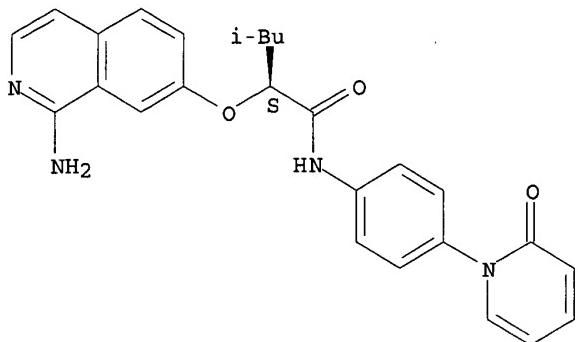
● HCl

09/ 830,227

RN 498541-33-6 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyloxy)-4-methyl-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]-, monohydrochloride, (2S)- (9CI) (CA INDEX NAME)

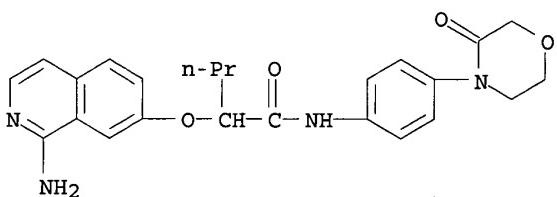
Absolute stereochemistry.



● HCl

RN 498541-35-8 CAPLUS

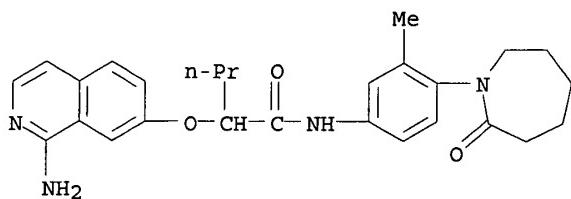
CN Pentanamide, 2-[(1-amino-7-isoquinolinyloxy)-N-[4-(3-oxo-4-morpholinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-37-0 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyloxy)-N-[4-(hexahydro-2-oxo-1H-azepin-1-yl)-3-methylphenyl]-, monohydrochloride (9CI) (CA INDEX NAME)

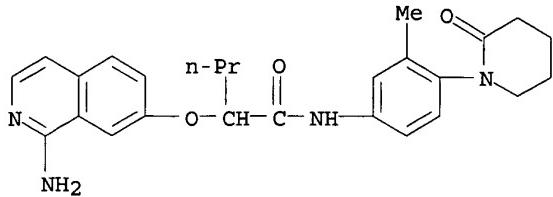


HCl

09/ 830,227

RN 498541-38-1 CAPLUS

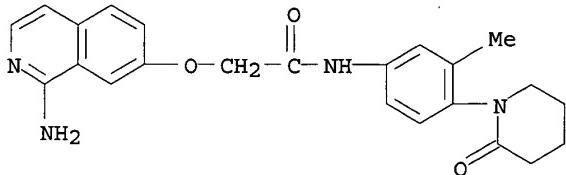
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-39-2 CAPLUS

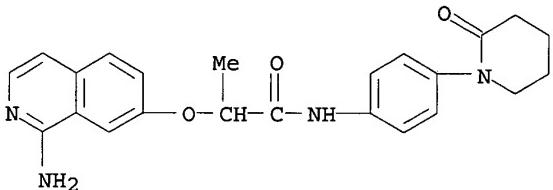
CN Acetamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-56-3 CAPLUS

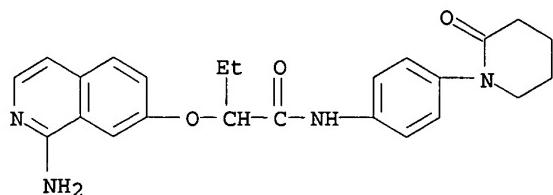
CN Propanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-58-5 CAPLUS

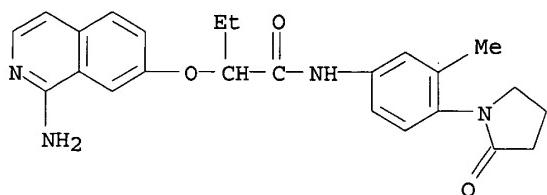
CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-60-9 CAPLUS

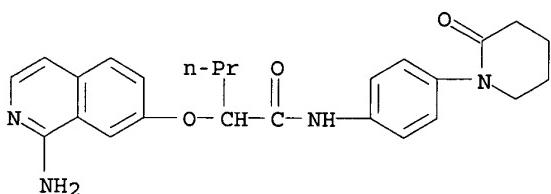
CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(2-oxo-1-pyrrolidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-62-1 CAPLUS

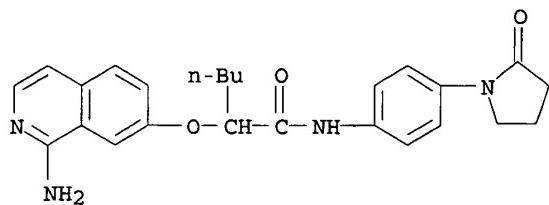
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-64-3 CAPLUS

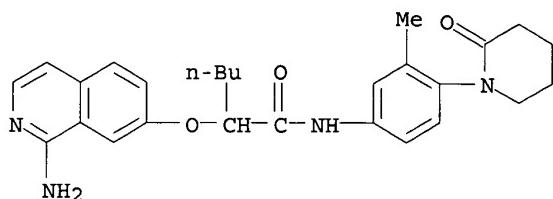
CN Hexanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1-pyrrolidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-66-5 CAPLUS

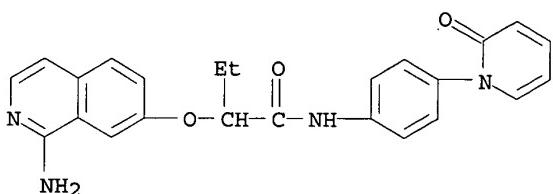
CN Hexanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-67-6 CAPLUS

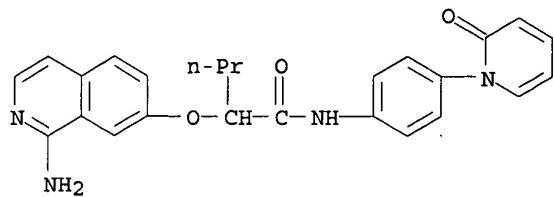
CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-68-7 CAPLUS

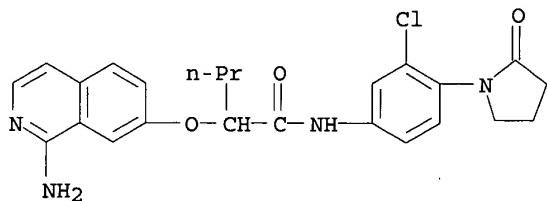
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-69-8 CAPLUS

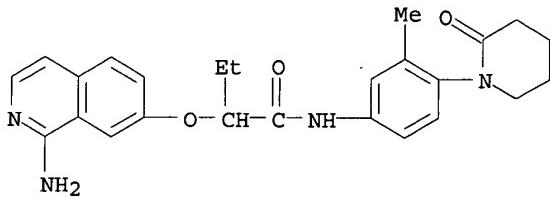
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-chloro-4-(2-oxo-1-pyrrolidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-70-1 CAPLUS

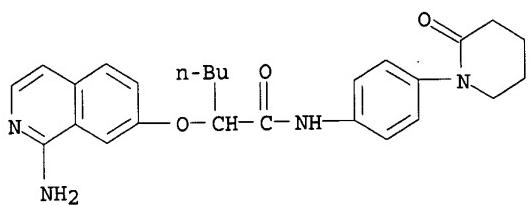
CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

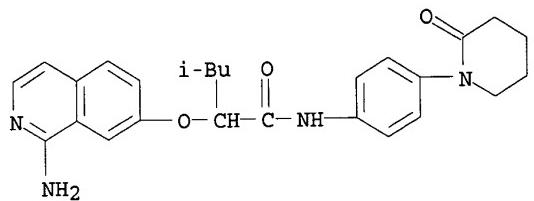
RN 498541-71-2 CAPLUS

CN Hexanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



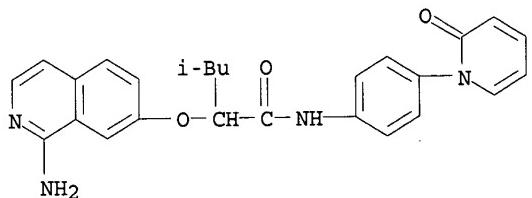
● HCl

RN 498541-72-3 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



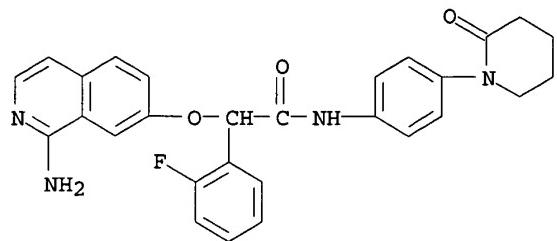
● HCl

RN 498541-73-4 CAPLUS
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

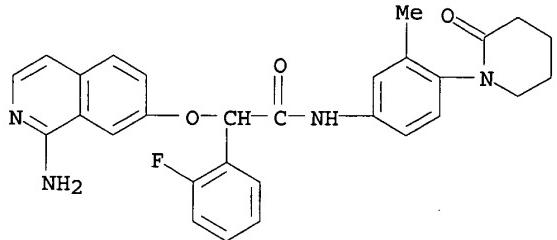
RN 498541-74-5 CAPLUS
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-2-fluoro-N-[4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-75-6 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-2-fluoro-N-[3-methyl-4-(2-oxo-1-piperidinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)

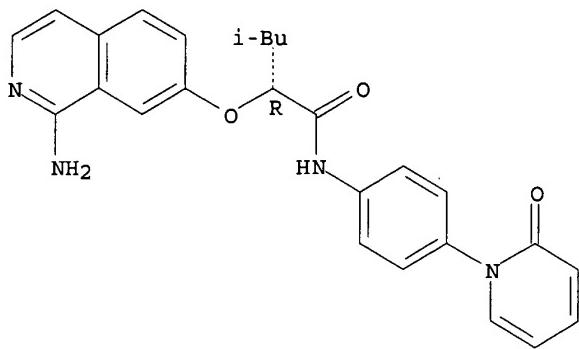


● HCl

RN 498541-76-7 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]-, monohydrochloride, (2R)- (9CI) (CA INDEX NAME)

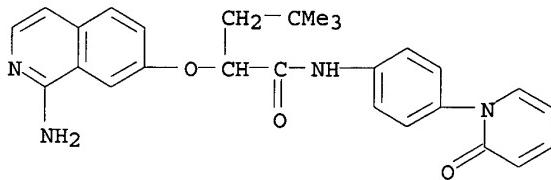
Absolute stereochemistry.



● HCl

RN 498541-78-9 CAPLUS

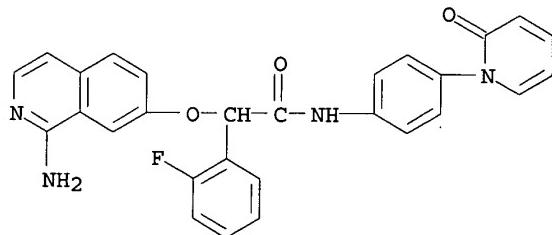
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4,4-dimethyl-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-80-3 CAPLUS

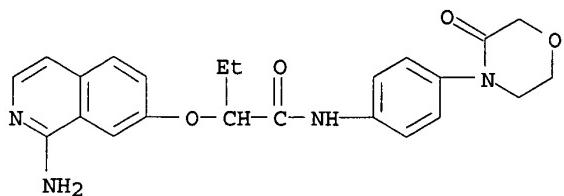
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-2-fluoro-N-[4-(2-oxo-1(2H)-pyridinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-82-5 CAPLUS

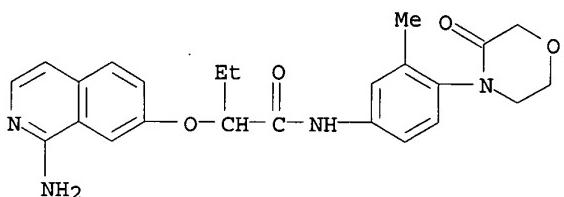
CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(3-oxo-4-morpholinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-84-7 CAPLUS

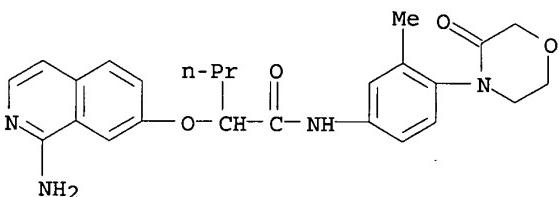
CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(3-oxo-4-morpholinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-87-0 CAPLUS

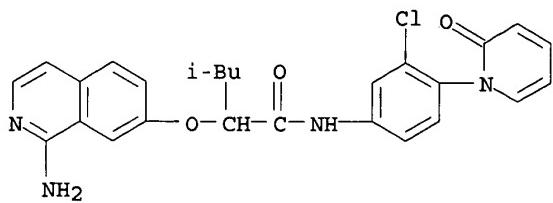
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-methyl-4-(3-oxo-4-morpholinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-88-1 CAPLUS

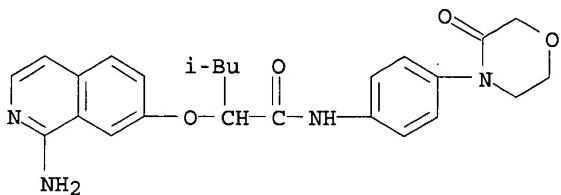
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[3-chloro-4-(2-oxo-1(2H)-pyridinyl)phenyl]-4-methyl-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-89-2 CAPLUS

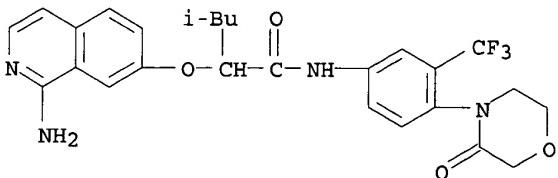
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(3-oxo-4-morpholinyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-90-5 CAPLUS

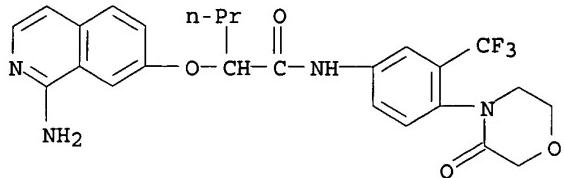
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[4-(3-oxo-4-morpholinyl)-3-(trifluoromethyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)



● HCl

RN 498541-92-7 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(3-oxo-4-morpholinyl)-3-(trifluoromethyl)phenyl]-, monohydrochloride (9CI) (CA INDEX NAME)

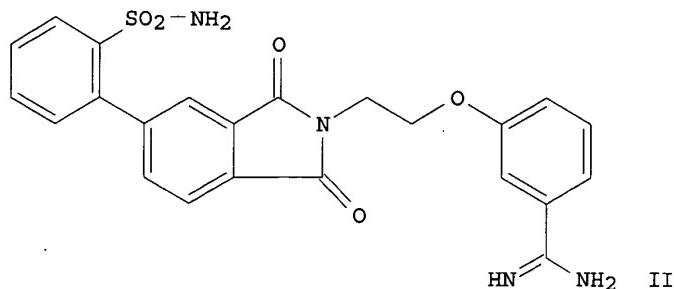
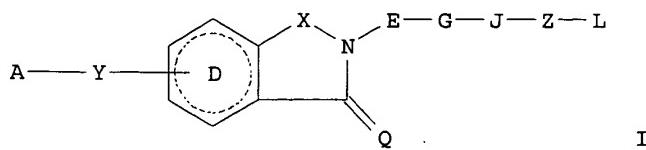


● HCl

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 3 OF 10 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 2002:927401 CAPLUS
 DOCUMENT NUMBER: 138:14016
 TITLE: Preparation of isoindole and isoquinoline derivatives as inhibitors of Factor xa
 INVENTOR(S): Zhang, Penglie; Zhu, Bing-Yan; Huang, Wenrong; Scarborough, Robert M.
 PATENT ASSIGNEE(S): Millennium Pharmaceuticals, Inc., USA
 SOURCE: PCT Int. Appl., 72 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--|------------------|-----------------|------------|
| WO 2002096873 | A1 | 20021205 | WO 2002-US16784 | 20020529 |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | |
| US 2003114448 | A1 | 20030619 | US 2002-171804 | 20020528 |
| PRIORITY APPLN. INFO.: | | | US 2001-294273P | P 20010531 |
| OTHER SOURCE(S): | | MARPAT 138:14016 | | |
| GI | | | | |



AB Isoindole and isoquinoline derivs. [I; wherein A = H, (C₁-C₆)alkyl, (C₃-C₈)cycloalkyl, alkylamino, alkenylamino, (substituted) Ph, etc.; Y = a bond, C(:O), CH₂, alkylamino, amide, etc.; D = (substituted) Ph, five- or six-membered arom. heterocyclic ring having from 1-2 hetero atoms selected from O, S, and N; X = alkylcarboxy, alkylsulfoxy, C(:O), C(:S), etc.; Q = O, or Q and the carbon atom to which it is attached is CH₂; E = a bond, alkyl, C(:O), etc.; G = O, alkoxy, amino, S, S(:O), S(:O)₂, etc.; J = O, S, amino, S(:O), S(:O)₂, etc.; Z = (substituted) Ph, naphthyl, monocyclic or fused bicyclic heterocyclic ring, etc.; L = H, CN, amido, amino, alkoxy, etc.] were prep'd. For example, II was prep'd. by a multistep synthetic procedure. The prep'd. compds. have activity against mammalian factor Xa and, thus, the compds. are useful in vitro or in vivo for preventing or treating coagulation disorders.

IT 476352-90-6P 476352-91-7P 476352-92-8P

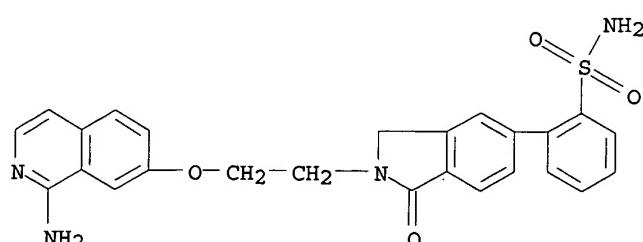
476352-93-9P

RL: PAC (Pharmacological activity); **SPN:** (Synthetic preparation); **THU:** (Therapeutic use); **BIOL:** (Biological study); **PREP:** (Preparation); **USES:** (Uses)

(prepn. of isoindole and isoquinoline derivs. as inhibitors of Factor Xa)

RN 476352-90-6 CAPLUS

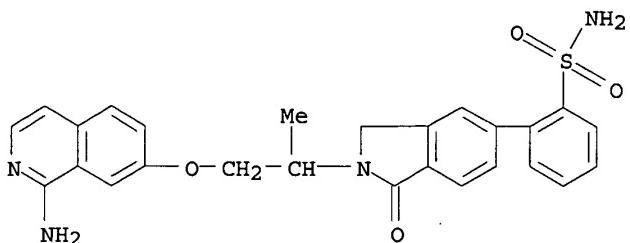
CN Benzenesulfonamide, 2-[2-[2-[(1-amino-7-isoquinolinyl)oxy]ethyl]-2,3-dihydro-1-oxo-1H-isoindol-5-yl]- (9CI) (CA INDEX NAME)



09/ 830,227

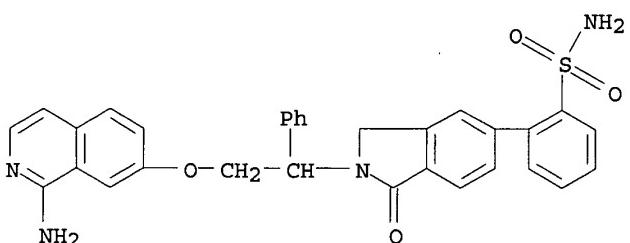
RN 476352-91-7 CAPLUS

CN Benzenesulfonamide, 2-[2-[2-[(1-amino-7-isoquinolinyloxy)-1-methylethyl]-2,3-dihydro-1-oxo-1H-isoindol-5-yl]- (9CI) (CA INDEX NAME)



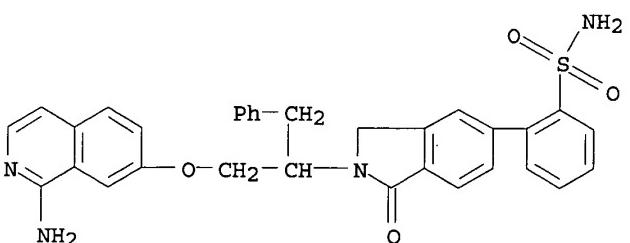
RN 476352-92-8 CAPLUS

CN Benzenesulfonamide, 2-[2-[2-[(1-amino-7-isoquinolinyloxy)-1-phenylethyl]-2,3-dihydro-1-oxo-1H-isoindol-5-yl]- (9CI) (CA INDEX NAME)



RN 476352-93-9 CAPLUS

CN Benzenesulfonamide, 2-[2-[1-[(1-amino-7-isoquinolinyloxy)methyl]-2-phenylethyl]-2,3-dihydro-1-oxo-1H-isoindol-5-yl]- (9CI) (CA INDEX NAME)



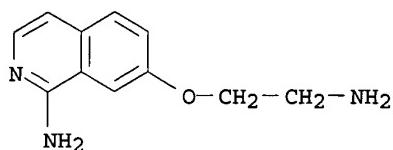
IT 309930-41-4 476352-88-2 476352-89-3

RL: RCT (Reactant); RACT (Reactant or reagent)

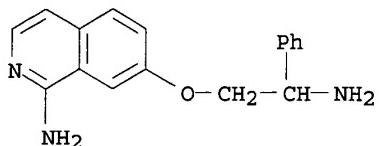
(prepn. of isoindole and isoquinoline derivs. as inhibitors of Factor xa)

RN 309930-41-4 CAPLUS

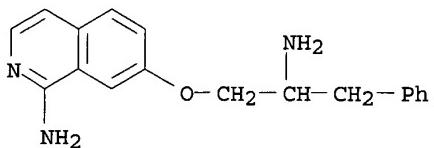
CN 1-Isoquinolinamine, 7-(2-aminoethoxy)- (9CI) (CA INDEX NAME)



RN 476352-88-2 CAPLUS
 CN 1-Isoquinolinamine, 7-(2-amino-2-phenylethoxy)- (9CI) (CA INDEX NAME)



RN 476352-89-3 CAPLUS
 CN 1-Isoquinolinamine, 7-(2-amino-3-phenylpropoxy)- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 4 OF 10 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 2002:407965 CAPLUS
 DOCUMENT NUMBER: 137:384703
 TITLE: Design, synthesis, and SAR of monobenzamidines and aminoisoquinolines as factor Xa inhibitors
 AUTHOR(S): Zhang, Penglie; Zuckett, Jingmei F.; Woolfrey, John;
 Tran, Katherine; Huang, Brian; Wong, Paul; Sinha, Uma;
 Park, Gary; Reed, Andrea; Malinowski, John;
 Hollenbach, Stan; Scarborough, Robert M.; Zhu,
 Bing-Yan
 CORPORATE SOURCE: Department of Medicinal Chemistry, Millennium
 Pharmaceuticals, Inc., South San Francisco, CA, 94080,
 USA
 SOURCE: Bioorganic & Medicinal Chemistry Letters (2002),
 12(12), 1657-1661
 CODEN: BMCLE8; ISSN: 0960-894X
 PUBLISHER: Elsevier Science Ltd.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Monoamidine FXa inhibitors, e.g. I (R = H, Me, Ph, PhCH₂), were designed

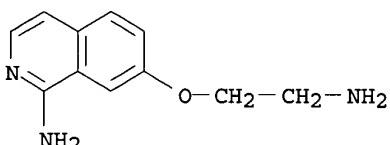
and synthesized. SAR studies and mol. modeling led to the design of conformationally constrained diaryl ethers, e.g. II [X = C(O)NH, NHCO], as well as benzopyrrolidinone III as potent FXa inhibitors. The monoamidines show high efficacy in a DVT model, but lack desirable oral bioavailability. The benzopyrrolidinone-based aminoisoquinolines, e.g. IV, do not show significant improvement in oral bioavailability.

IT 309930-41-4P 476352-87-1P 476352-88-2P
476352-89-3P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(lactamization; prepn. of phenyl(oxoisoindoline)ethoxy(isoquinolinamine) as factor Xa inhibitors)

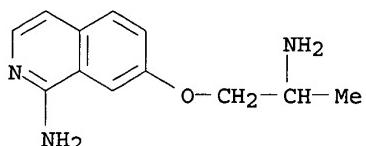
RN 309930-41-4 CAPLUS

CN 1-Isoquinolinamine, 7-(2-aminoethoxy)- (9CI) (CA INDEX NAME)



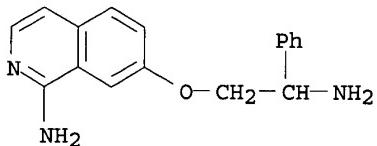
RN 476352-87-1 CAPLUS

CN 1-Isoquinolinamine, 7-(2-aminopropoxy)- (9CI) (CA INDEX NAME)



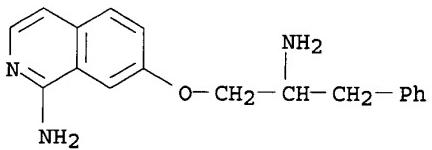
RN 476352-88-2 CAPLUS

CN 1-Isoquinolinamine, 7-(2-amino-2-phenylethoxy)- (9CI) (CA INDEX NAME)



RN 476352-89-3 CAPLUS

CN 1-Isoquinolinamine, 7-(2-amino-3-phenylpropoxy)- (9CI) (CA INDEX NAME)



IT 476352-90-6P 476352-91-7P 476352-92-8P

476352-93-9P

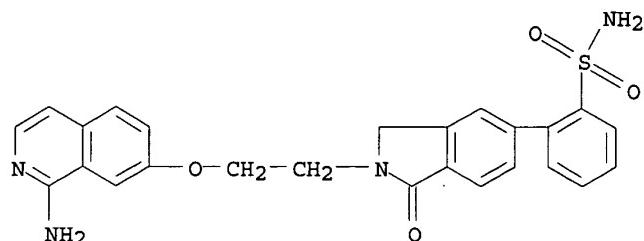
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); BIOL

09/ 830,227

(Biological study); PREP (Preparation)
(prep. of phenyl(oxoisoindoline)ethoxy(isoquinolinamine) as factor Xa
inhibitors)

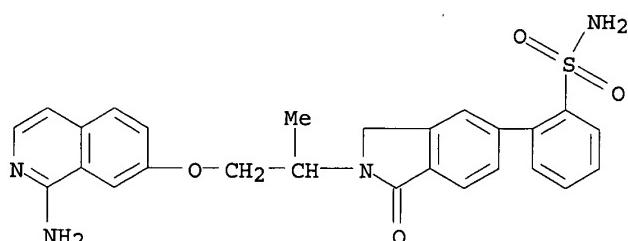
RN 476352-90-6 CAPLUS

CN Benzenesulfonamide, 2-[2-[2-[(1-amino-7-isoquinolinyloxy)ethyl]-2,3-dihydro-1-oxo-1H-isoindol-5-yl]- (9CI) (CA INDEX NAME)



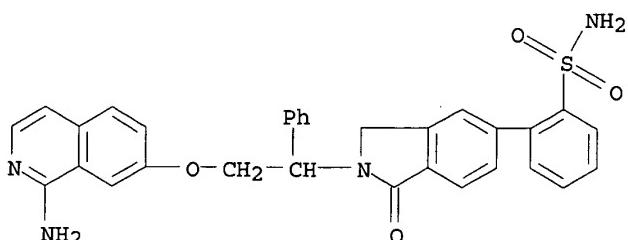
RN 476352-91-7 CAPLUS

CN Benzenesulfonamide, 2-[2-[2-[(1-amino-7-isoquinolinyloxy)-1-methylethyl]-2,3-dihydro-1-oxo-1H-isoindol-5-yl]- (9CI) (CA INDEX NAME)



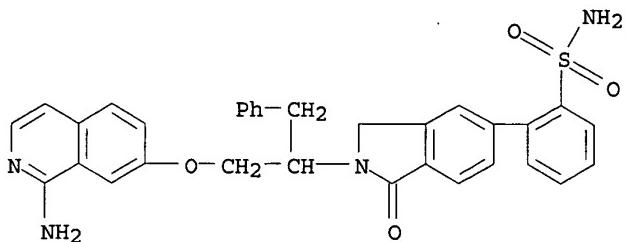
RN 476352-92-8 CAPLUS

CN Benzenesulfonamide, 2-[2-[2-[(1-amino-7-isoquinolinyloxy)-1-phenylethyl]-2,3-dihydro-1-oxo-1H-isoindol-5-yl]- (9CI) (CA INDEX NAME)



RN 476352-93-9 CAPLUS

CN Benzenesulfonamide, 2-[2-[1-[(1-amino-7-isoquinolinyloxy)methyl]-2-phenylethyl]-2,3-dihydro-1-oxo-1H-isoindol-5-yl]- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 5 OF 10 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2002:240733 CAPLUS

DOCUMENT NUMBER: 136:263103

TITLE: Biphenyl-substituted aminoquinolines and -isoquinolines as factor Xa inhibitors

INVENTOR(S): Dorsch, Dieter; Juraszyk, Horst; Mederski, Werner; Tsaklakidis, Christos; Gleitz, Johannes; Barnes, Christopher

PATENT ASSIGNEE(S): Merck Patent G.m.b.H., Germany

SOURCE: PCT Int. Appl., 36 pp.

CODEN: PIXXD2

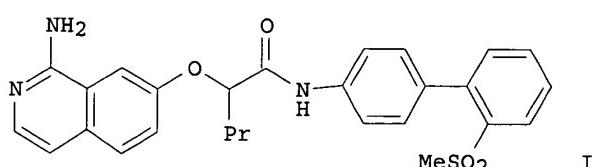
DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|--------|------------|--------------------|----------|
| WO 2002024654 | A1 | 20020328 | WO 2001-EP10786 | 20010918 |
| W: CA, JP, US | | | | |
| RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR | | | | |
| DE 10046272 | A1 | 20020328 | DE 2000-10046272 | 20000919 |
| EP 1322618 | A1 | 20030702 | EP 2001-985251 | 20010918 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | | |
| PRIORITY APPLN. INFO.: | | | DE 2000-10046272 A | 20000919 |
| | | | WO 2001-EP10786 W | 20010918 |
| OTHER SOURCE(S): | MARPAT | 136:263103 | | |
| GI | | | | |



I

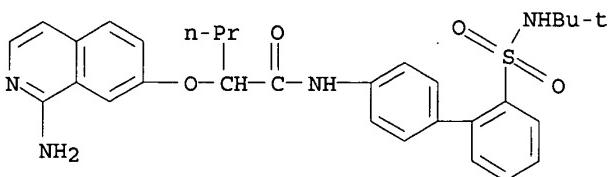
AB The title compds. were prep'd. for use as inhibitors of blood coagulation factors Xa and VIIa (no data). Thus, 7-isoquinolinol was treated with BrCHPrCO2CMe3, followed by ester hydrolysis, amidation with 2-MeSO2C6H4C6H4NH2-4, N-oxidn., reaction with pyridine, and treatment with ethanolamine to give the title compd. I.

IT 405272-07-3

RL: RCT (Reactant); RACT (Reactant or reagent)
 (prepn. of biphenyl-substituted aminoquinolines and -isoquinolines as
 factor Xa inhibitors)

RN 405272-07-3 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-[[[(1,1'-dimethylethyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)

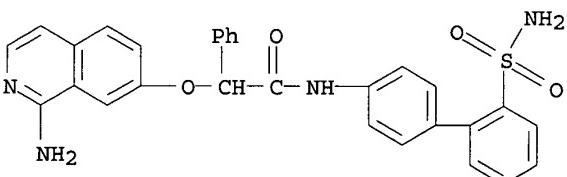


IT 308288-71-3P 405272-04-0P 405272-05-1P
 405272-06-2P 405272-08-4P 405272-09-5P
 405272-10-8P 405272-11-9P 405272-12-0P
 405272-13-1P 405272-14-2P 405272-17-5P
 405272-18-6P 405272-19-7P 405272-20-0P
 405272-21-1P 405272-22-2P 405272-23-3P
 405272-24-4P 405272-25-5P 405272-26-6P

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (prepn. of biphenyl-substituted aminoquinolines and -isoquinolines as factor Xa inhibitors)

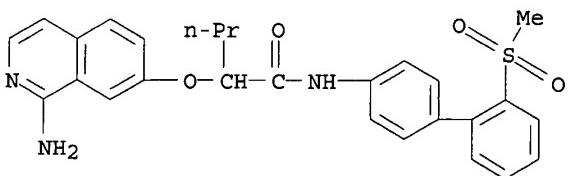
RN 308288-71-3 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



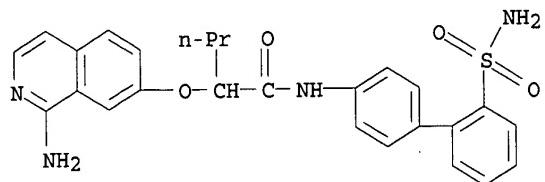
RN 405272-04-0 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(methylsulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



RN 405272-05-1 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



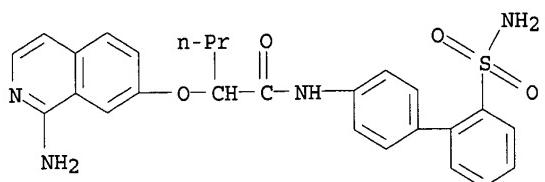
RN 405272-06-2 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]-, trifluoroacetate (9CI) (CA INDEX NAME)

CM 1

CRN 405272-05-1

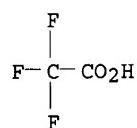
CMF C26 H26 N4 O4 S



CM 2

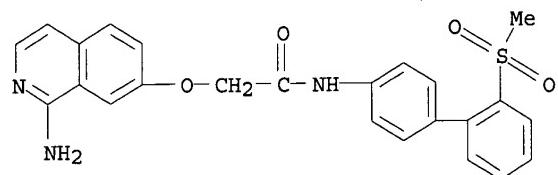
CRN 76-05-1

CMF C2 H F3 O2



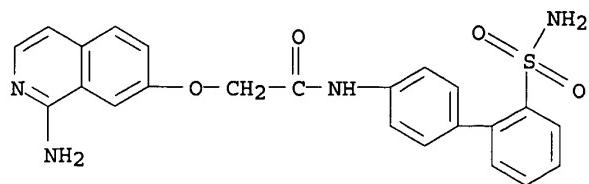
RN 405272-08-4 CAPLUS

CN Acetamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(methylsulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



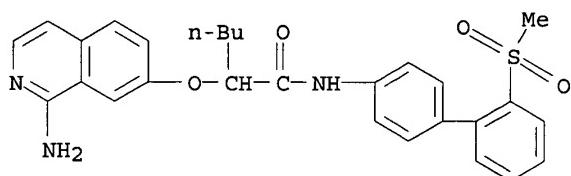
RN 405272-09-5 CAPLUS

CN Acetamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



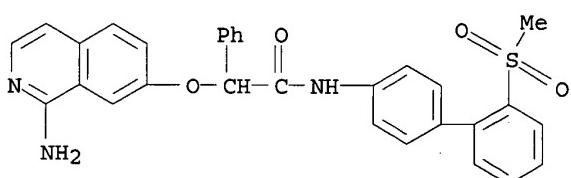
RN 405272-10-8 CAPLUS

CN Hexanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(methylsulfonyl)[1,1'-biphenyl]-4-yl] - (9CI) (CA INDEX NAME)



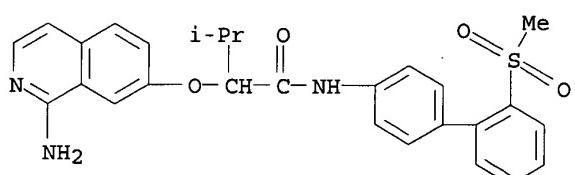
RN 405272-11-9 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(methylsulfonyl)[1,1'-biphenyl]-4-yl] - (9CI) (CA INDEX NAME)



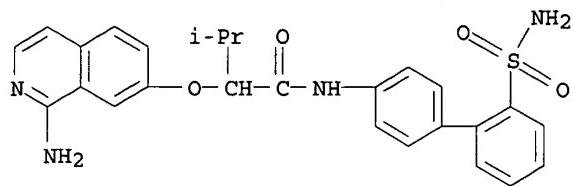
RN 405272-12-0 CAPLUS

CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-3-methyl-N-[2'-(methylsulfonyl)[1,1'-biphenyl]-4-yl] - (9CI) (CA INDEX NAME)



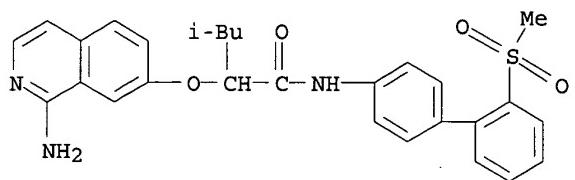
RN 405272-13-1 CAPLUS

CN Butanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]-3-methyl- (9CI) (CA INDEX NAME)



RN 405272-14-2 CAPLUS

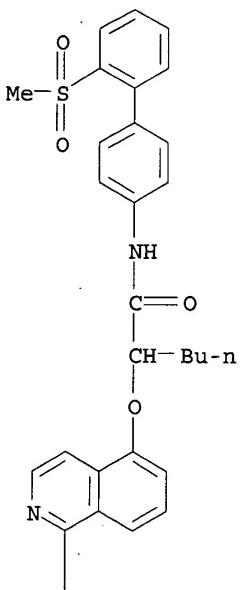
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-4-methyl-N-[2'-(methylsulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



RN 405272-17-5 CAPLUS

CN Hexanamide, 2-[(1-amino-5-isoquinolinyl)oxy]-N-[2'-(methylsulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A

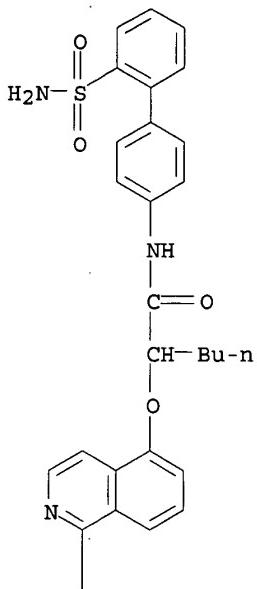


09/ 830,227

RN 405272-18-6 CAPLUS

CN Hexanamide, 2-[(1-amino-5-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]-(9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A

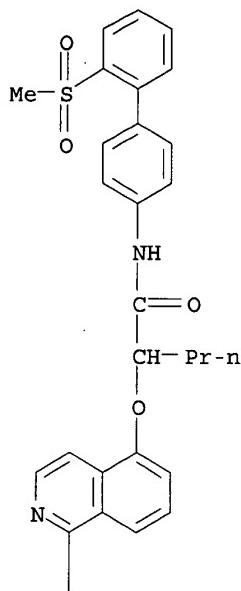


RN 405272-19-7 CAPLUS

CN Pentanamide, 2-[(1-amino-5-isoquinolinyl)oxy]-N-[2'-(methylsulfonyl)[1,1'-biphenyl]-4-yl]-(9CI) (CA INDEX NAME)

09/ 830,227

PAGE 1-A

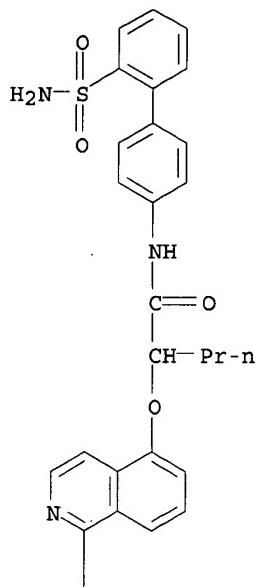


PAGE 2-A



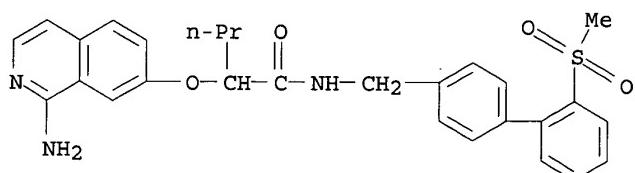
RN 405272-20-0 CAPLUS

CN Pentanamide, 2-[(1-amino-5-isoquinolinyloxy)-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



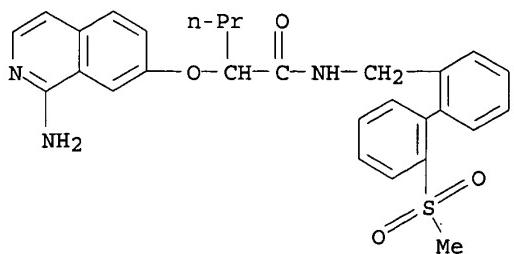
RN 405272-21-1 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[[2'-(methylsulfonyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)



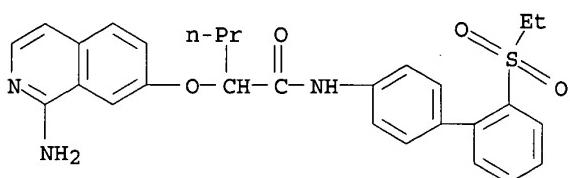
RN 405272-22-2 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[[2'-(methylsulfonyl)[1,1'-biphenyl]-2-yl]methyl]- (9CI) (CA INDEX NAME)



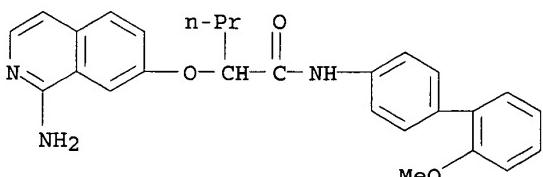
RN 405272-23-3 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(ethylsulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



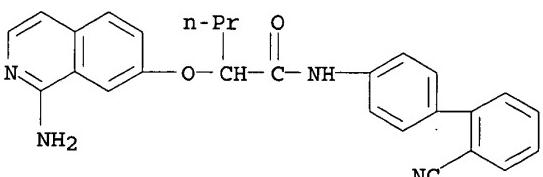
RN 405272-24-4 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-(2'-methoxy[1,1'-biphenyl]-4-yl)- (9CI) (CA INDEX NAME)



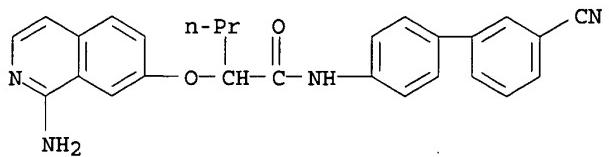
RN 405272-25-5 CAPLUS

CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-(2'-cyano[1,1'-biphenyl]-4-yl)- (9CI) (CA INDEX NAME)



RN 405272-26-6 CAPLUS

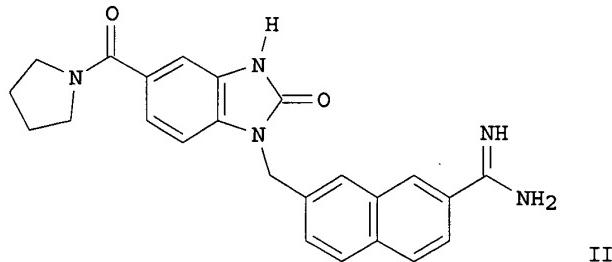
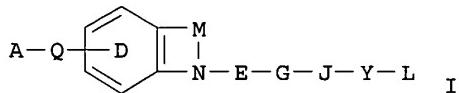
CN Pentanamide, 2-[(1-amino-7-isoquinolinyl)oxy]-N-(3'-cyano[1,1'-biphenyl]-4-yl)- (9CI) (CA INDEX NAME)



REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 6 OF 10 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 2001:137189 CAPLUS
 DOCUMENT NUMBER: 134:193446
 TITLE: Preparation of heterocyclic compounds as inhibitors of factor Xa
 INVENTOR(S): Zhu, Bing-Yan; Scarborough, Robert M.; Clizbe, Lane;
 Doughan, Brandon; Jia, Zhaozhong-Jon; Kane-Maguire,
 Kim, Marlowe, Charles; Song, Yonghong; Su, Ting; Teng,
 Willy; Zhang, Penglie
 PATENT ASSIGNEE(S): Cor Therapeutics, Inc., USA; et al.
 SOURCE: PCT Int. Appl., 387 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--|-------------------|-----------------|------------|
| WO 2001012600 | A1 | 20010222 | WO 2000-US21742 | 20000810 |
| WO 2001012600 | C2 | 20020912 | | |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |
| US 6534535 | B1 | 20030318 | US 2000-636804 | 20000810 |
| PRIORITY APPLN. INFO.: | | | US 1999-148627P | P 19990812 |
| | | | US 2000-202202P | P 20000505 |
| OTHER SOURCE(S): GI | | MARPAT 134:193446 | | |



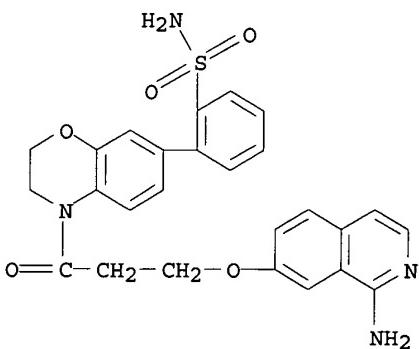
AB The title compds. [I; A = alkyl, cycloalkyl, (un)substituted Ph, etc.; Q = a direct link, CH₂, CO, etc.; D = (un)substituted Ph, 6-membered heteroaryl having 1-2 ring N atoms; M = NR₁₆CO, NR₁₆CS, CR₁₇R₁₈CO, etc.; R₁₆-R₁₈ = H, halo, alkyl, etc.; E = a direct link, CO, CONR₅, etc.; R₅ = alkyl, alkenyl, alkynyl, etc.; G = a direct link, CR₇R₈, CR₇aR₈aCR₇bR₈b, CR₇c:CR₈c; R₇, R₈, R₇a, R₇b, R₇c, R₈a, R₈b, R₈c = H, halo, alkyl, etc.; J = a direct link, O, S, etc.; Y = (un)substituted Ph, naphthyl, monocyclic or fused bicyclic heterocyclyl; L = H, CN, CONR₁₂R₁₃; R₁₂, R₁₃ = H, alkyl, OH, etc.] having activity against mammalian factor Xa, and useful in vitro or in vivo for preventing or treating coagulation disorders, were prepd. and formulated. E.g., a multi-step synthesis of the title compd. II was given.

IT **327046-29-7P**

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(prepn. of heterocyclic compds. as inhibitors of factor Xa)

RN 327046-29-7 CAPLUS

CN 2H-1,4-Benzoxazine, 4-[3-[(1-amino-7-isouquinolinyl)oxy]-1-oxopropyl]-7-[2-(aminosulfonyl)phenyl]-3,4-dihydro- (9CI) (CA INDEX NAME)



REFERENCE COUNT:

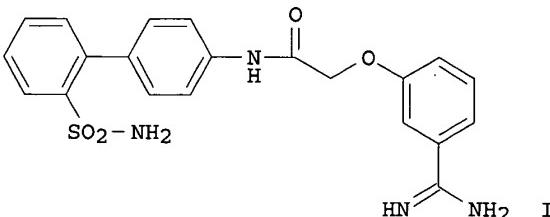
13

THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ACCESSION NUMBER: 2000:842106 CAPLUS
 DOCUMENT NUMBER: 134:29205
 TITLE: Preparation of benzamidines and arylamidines as inhibitors of factor Xa
 INVENTOR(S): Su, Ting; Zhu, Bing-Yan; Kane-Maguire, Kim; Scarborough, Robert M.; Zhang, Penglie
 PATENT ASSIGNEE(S): Cor Therapeutics, Inc., USA
 SOURCE: PCT Int. Appl., 144 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 3
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--|----------|-------------------|----------|
| WO 2000071510 | A2 | 20001130 | WO 2000-US14195 | 20000524 |
| WO 2000071510 | A3 | 20010830 | | |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |
| EP 1183235 | A2 | 20020306 | EP 2000-937700 | 20000524 |
| R: | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | |
| JP 2003500385 | T2 | 20030107 | JP 2000-619767 | 20000524 |
| PRIORITY APPLN. INFO.: | | | US 1999-135849P P | 19990524 |
| | | | WO 2000-US14195 W | 20000524 |

OTHER SOURCE(S): MARPAT 134:29205
 GI



AB AYDEGJZL [wherein A = (cyclo)alkyl, NR₂R₃, C(:NR₂)NR₂R₃, C(:NR₂)R₃, NR₃C(:NR₂)NR₂NR₃, (un)substituted Ph, naphthyl, or heterocyclic ring; R₂ and R₃ = independently H, (cyclo)alkyl, alkenyl, alkynyl, alkylcyclalkyl, or (un)substituted alkylphenyl or alkynlnaphthyl; Y = bond, bivalent alkyl, alkenyl, or alkynyl, CH₂, CO, C(:NR₄), NR₄, NR₄CH₂, CH₂NR₄, CONR₄, NR₄CO, SO₂, O, SO₂NR₄, or NR₄SO₂; R₄ = H, alkyl, alkenyl, alkynyl, or (un)substituted aklylaryl or aklyheterocyclyl; D = (un)substituted Ph, naphthyl, or heterocyclic ring; E = NR₅CO, CONR₅, NR₅, or NR₅(CH₂)₀₋₂; R₅ = H, alkyl, alkyl(hetero)aryl, or (un)substituted carboxyaklyl or carboxamidoaklyl; G = (un)substituted methylene or ethylene; J = O, OCHR₁₁, S, SCHR₁₁, S(O), SO₂, S(O)CHR₁₁, SO₂CHR₁₁; R₁₁ = H, alkyl, or (un)substituted alkyl(hetero)aryl; Z = (un)substituted Ph, naphthyl, or

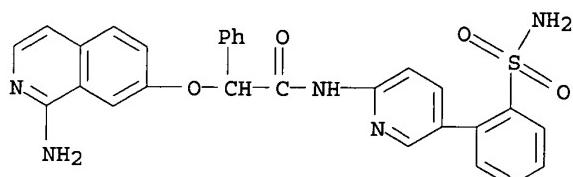
heterocyclic ring; L = H, CN, CONR12NR13, (CH₂)₀₋₂NR12R13, C(:NR12)NR12R13, NR12R13, OR12, NR12C(:NR12)NR12N13, or NR12C(:N12)R13; R12 and R13 = independently H, OR14, NR14R15, alkyl, (un)substituted alkylphenyl, alkynaphthyl, or carboxyalkyl; R14 and R15 = independently H, alkyl, (un)substituted alkyl(hetero)aryl, or together with the attached N forms a heterocyclic ring] were prep'd. as potent and highly selective inhibitors of factor Xa for the prevention or treatment of coagulation disorders (no data). For example, 2-(3-cyanophenoxy)acetic acid was coupled with {[2-(4-aminophenyl)phenyl]sulfonyl}(tert-butyl)amine in the presence of BOP in DMF to give the acetamide intermediate. Treatment with NH₂OH.bul.HCl and TEA in EtOH, followed by addn. of AcOH, redn. using Pd/C in MeOH, and deprotection with TFA afforded the benzamidine (I). Compds. of the invention show selectivity for factor Xa vs. other proteases of the coagulation cascade or the fibrinolytic cascade, and are useful as diagnostic reagents as well as antithrombotic agents (no data).

IT 489426-93-9 489426-94-0 489426-96-2
 489426-98-4 489427-05-6 489427-06-7
 489427-07-8 489427-10-3 489427-11-4
 489427-15-8 489427-20-5 489427-40-9
 489427-42-1 489427-44-3 489427-50-1
 489427-54-5 489427-55-6 489427-57-8
 489427-59-0 489428-90-2 489428-91-3
 489428-92-4 489428-93-5 489428-94-6
 489428-95-7 489428-96-8 489428-97-9
 489428-98-0 489429-15-4 489429-16-5
 489429-17-6 489429-18-7 489429-19-8
 489429-22-3 489429-23-4 489429-24-5
 489429-31-4 489429-42-7 489429-45-0
 489429-63-2 489433-05-8 489434-39-1
 489438-63-3 489438-99-5 489448-10-4
 489448-25-1 489448-31-9 489448-49-9
 489448-64-8 489448-65-9 489448-66-0
 489448-67-1 308288-71-3P 308288-72-4P
 308288-75-7P 308288-76-8P 308288-77-9P
 308288-78-0P 308288-79-1P 308288-80-4P
 308288-83-7P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use)
 (prepn. of benzamidine and arylamidine factor Xa inhibitors by
 amidation of cyanoaryl-substituted carboxylic acids with amines and
 subsequent conversion of nitriles to amidines)

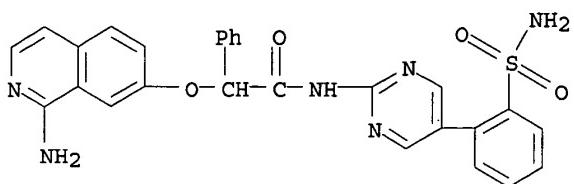
RN 489426-93-9 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[5-[2-(aminosulfonyl)phenyl]-2-pyridinyl]- (9CI) (CA INDEX NAME)



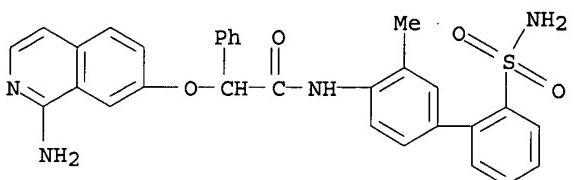
RN 489426-94-0 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[5-[2-(aminosulfonyl)phenyl]-2-pyrimidinyl]- (9CI) (CA INDEX NAME)



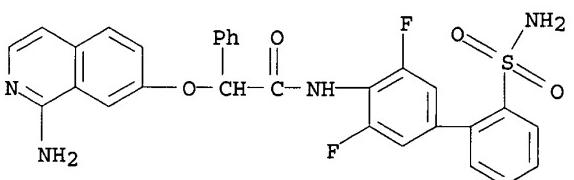
RN 489426-96-2 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)-3-methyl[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



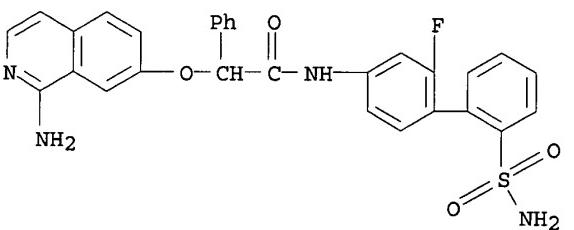
RN 489426-98-4 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)-3,5-difluoro[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



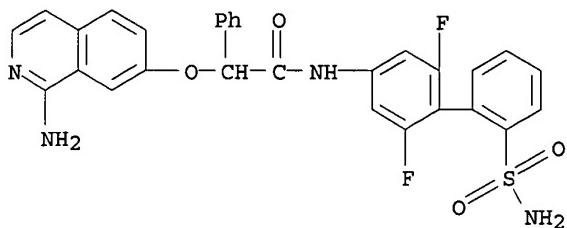
RN 489427-05-6 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)-2-fluoro[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



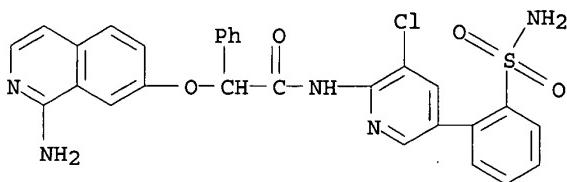
RN 489427-06-7 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)-2,6-difluoro[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



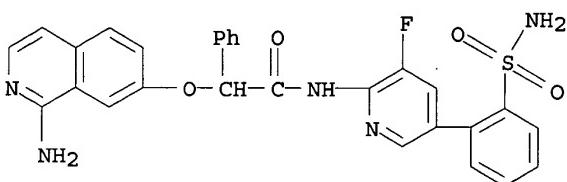
RN 489427-07-8 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[5-[2-(aminosulfonyl)phenyl]-3-chloro-2-pyridinyl]- (9CI) (CA INDEX NAME)



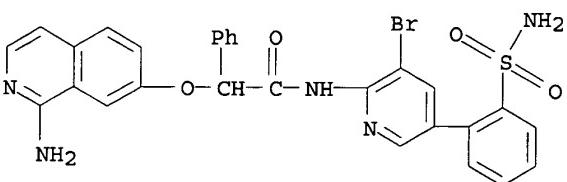
RN 489427-10-3 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[5-[2-(aminosulfonyl)phenyl]-3-fluoro-2-pyridinyl]- (9CI) (CA INDEX NAME)



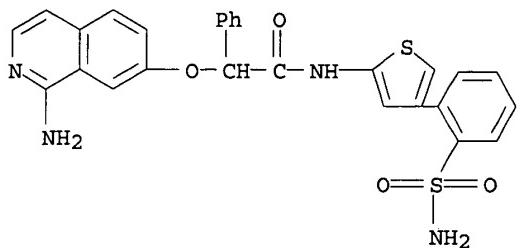
RN 489427-11-4 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[5-[2-(aminosulfonyl)phenyl]-3-bromo-2-pyridinyl]- (9CI) (CA INDEX NAME)



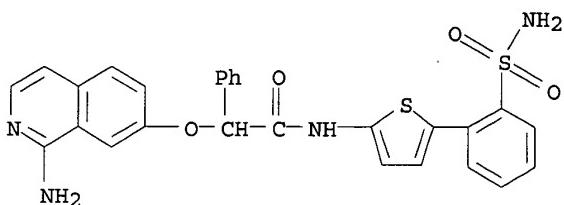
RN 489427-15-8 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[4-[2-(aminosulfonyl)phenyl]-2-thienyl]- (9CI) (CA INDEX NAME)



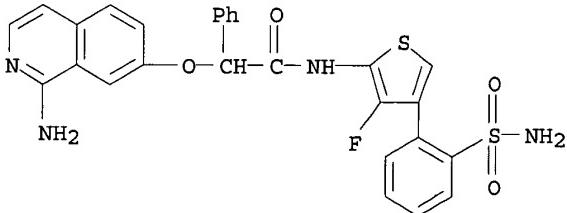
RN 489427-20-5 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[5-[2-(aminosulfonyl)phenyl]-2-thienyl]- (9CI) (CA INDEX NAME)



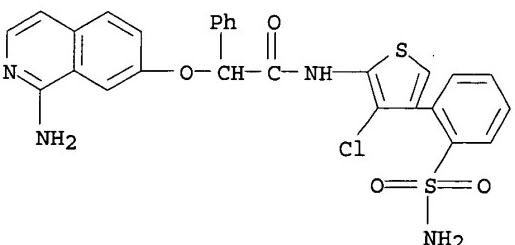
RN 489427-40-9 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[4-[2-(aminosulfonyl)phenyl]-3-fluoro-2-thienyl]- (9CI) (CA INDEX NAME)



RN 489427-42-1 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[4-[2-(aminosulfonyl)phenyl]-3-chloro-2-thienyl]- (9CI) (CA INDEX NAME)

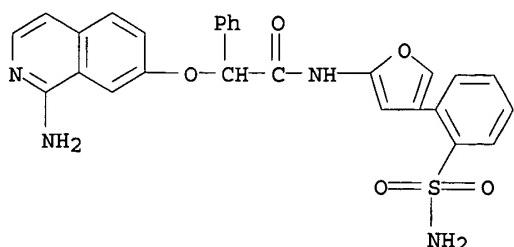


RN 489427-44-3 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[4-[2-

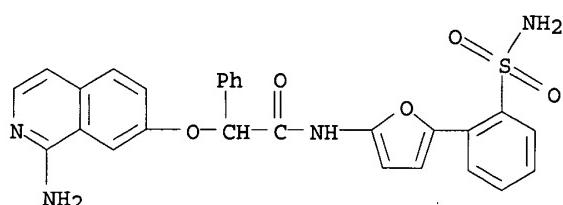
09/ 830,227

(aminosulfonyl)phenyl]-2-furanyl]- (9CI) (CA INDEX NAME)



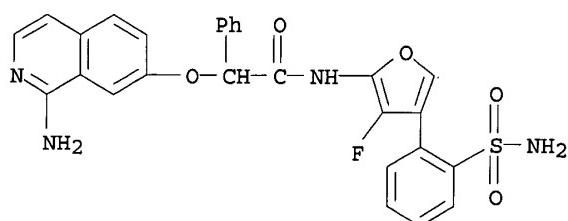
RN 489427-50-1 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[5-[2-(aminosulfonyl)phenyl]-2-furanyl]- (9CI) (CA INDEX NAME)



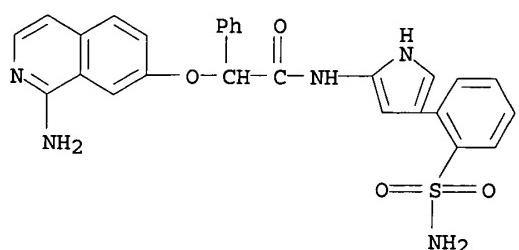
RN 489427-54-5 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[4-[2-(aminosulfonyl)phenyl]-3-fluoro-2-furanyl]- (9CI) (CA INDEX NAME)



RN 489427-55-6 CAPLUS

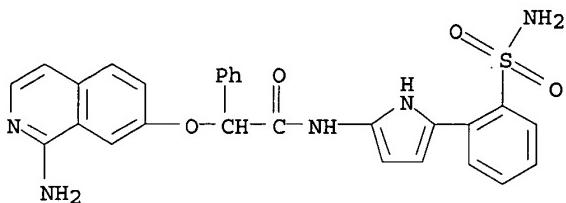
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[4-[2-(aminosulfonyl)phenyl]-1H-pyrrol-2-yl]- (9CI) (CA INDEX NAME)



09/ 830,227

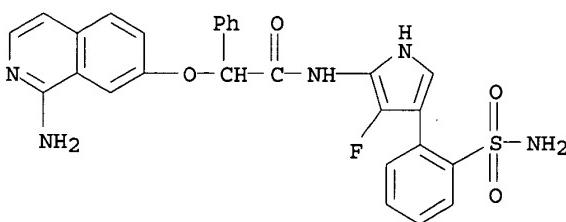
RN 489427-57-8 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[5-[2-(aminosulfonyl)phenyl]-1H-pyrrol-2-yl]- (9CI) (CA INDEX NAME)



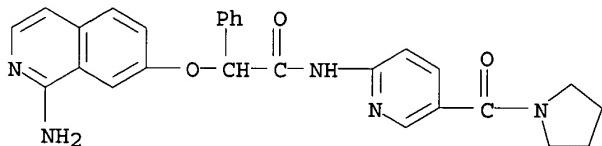
RN 489427-59-0 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[4-[2-(aminosulfonyl)phenyl]-3-fluoro-1H-pyrrol-2-yl]- (9CI) (CA INDEX NAME)



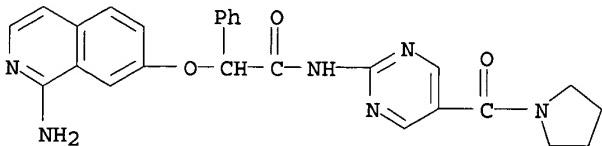
RN 489428-90-2 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[5-(1-pyrrolidinylcarbonyl)-2-pyridinyl]- (9CI) (CA INDEX NAME)



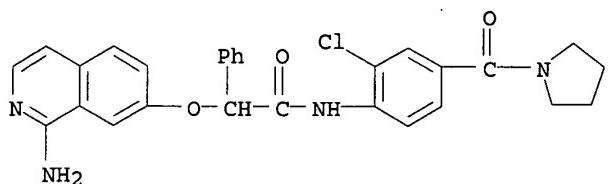
RN 489428-91-3 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[5-(1-pyrrolidinylcarbonyl)-2-pyrimidinyl]- (9CI) (CA INDEX NAME)



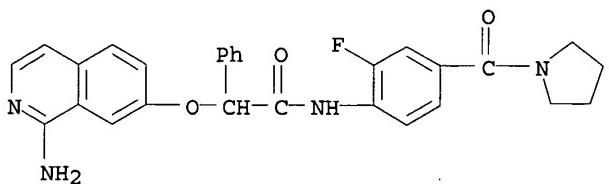
RN 489428-92-4 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[2-chloro-4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



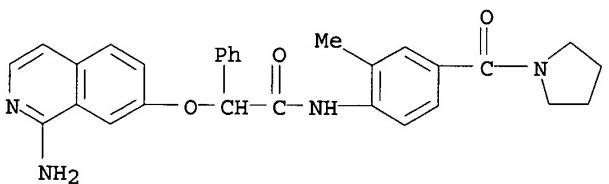
RN 489428-93-5 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2-fluoro-4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



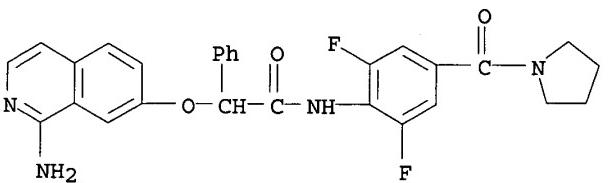
RN 489428-94-6 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2-methyl-4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



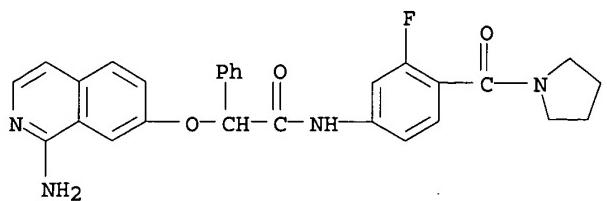
RN 489428-95-7 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isouquinolinyl)oxy]-N-[2,6-difluoro-4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)

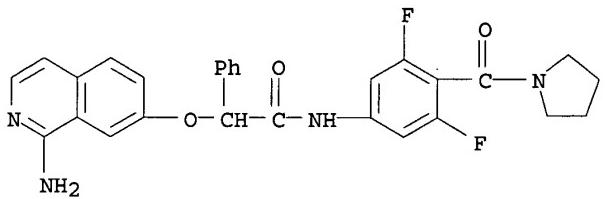


RN 489428-96-8 CAPLUS

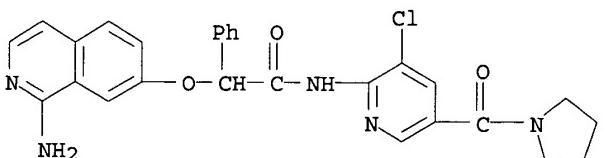
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[3-fluoro-4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



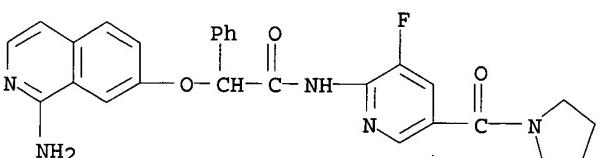
RN 489428-97-9 CAPLUS
CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[3,5-difluoro-4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



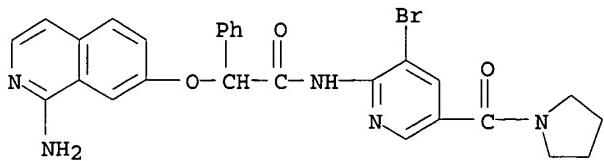
RN 489428-98-0 CAPLUS
CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[3-chloro-5-(1-pyrrolidinylcarbonyl)-2-pyridinyl]- (9CI) (CA INDEX NAME)



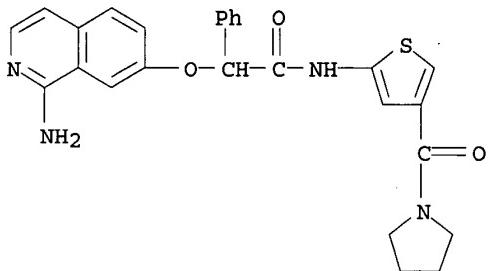
RN 489429-15-4 CAPLUS
CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[3-fluoro-5-(1-pyrrolidinylcarbonyl)-2-pyridinyl]- (9CI) (CA INDEX NAME)



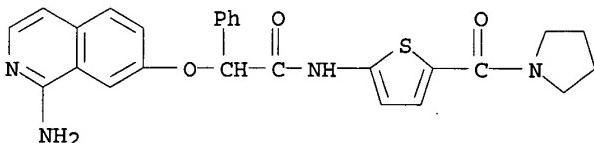
RN 489429-16-5 CAPLUS
CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[3-bromo-5-(1-pyrrolidinylcarbonyl)-2-pyridinyl]- (9CI) (CA INDEX NAME)



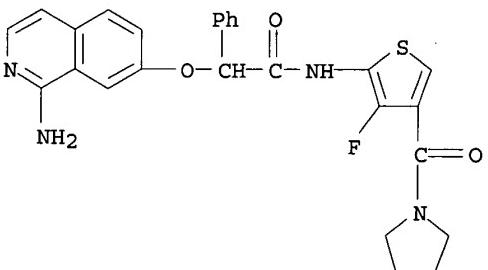
RN 489429-17-6 CAPLUS
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(1-pyrrolidinylcarbonyl)-2-thienyl]- (9CI) (CA INDEX NAME)



RN 489429-18-7 CAPLUS
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[5-(1-pyrrolidinylcarbonyl)-2-thienyl]- (9CI) (CA INDEX NAME)

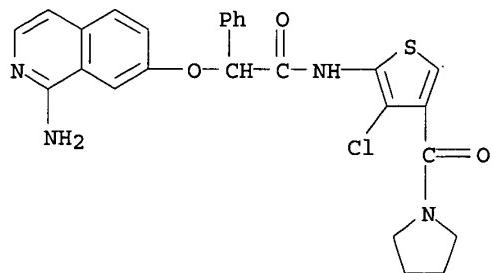


RN 489429-19-8 CAPLUS
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[3-fluoro-4-(1-pyrrolidinylcarbonyl)-2-thienyl]- (9CI) (CA INDEX NAME)



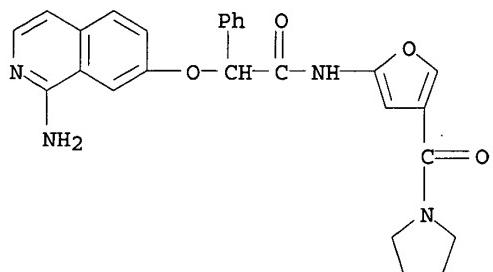
RN 489429-22-3 CAPLUS
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[3-chloro-4-(1-pyrrolidinylcarbonyl)-2-thienyl]- (9CI) (CA INDEX NAME)

09/ 830,227



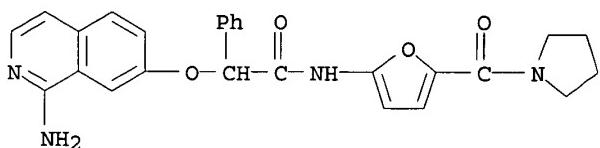
RN 489429-23-4 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(1-pyrrolidinylcarbonyl)-2-furanyl]- (9CI) (CA INDEX NAME)



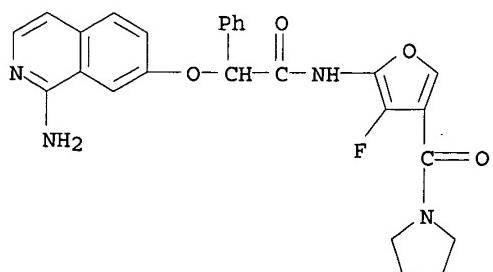
RN 489429-24-5 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[5-(1-pyrrolidinylcarbonyl)-2-furanyl]- (9CI) (CA INDEX NAME)



RN 489429-31-4 CAPLUS

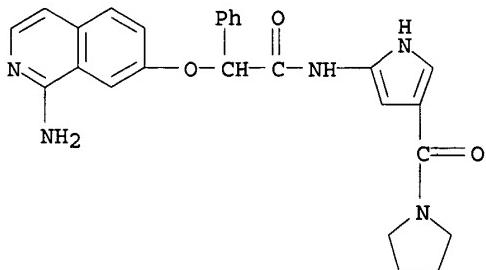
CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[3-fluoro-4-(1-pyrrolidinylcarbonyl)-2-furanyl]- (9CI) (CA INDEX NAME)



RN 489429-42-7 CAPLUS

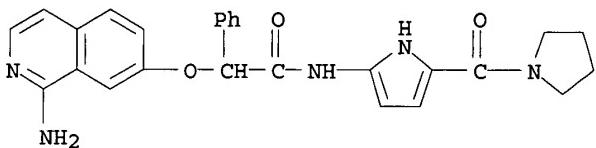
09/ 830,227

CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[4-(1-pyrrolidinylcarbonyl)-1H-pyrrol-2-yl]- (9CI) (CA INDEX NAME)



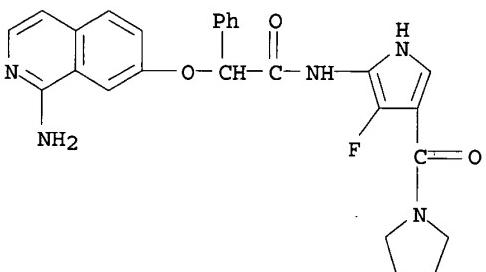
RN 489429-45-0 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[5-(1-pyrrolidinylcarbonyl)-1H-pyrrol-2-yl]- (9CI) (CA INDEX NAME)



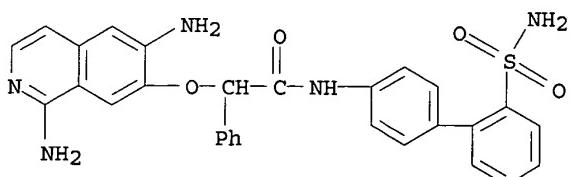
RN 489429-63-2 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[3-fluoro-4-(1-pyrrolidinylcarbonyl)-1H-pyrrol-2-yl]- (9CI) (CA INDEX NAME)



RN 489433-05-8 CAPLUS

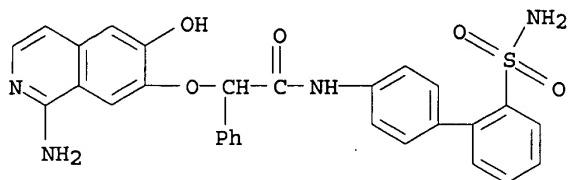
CN Benzeneacetamide, N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]-.alpha.-[(1,6-diamino-7-isooquinolinyl)oxy]- (9CI) (CA INDEX NAME)



RN 489434-39-1 CAPLUS

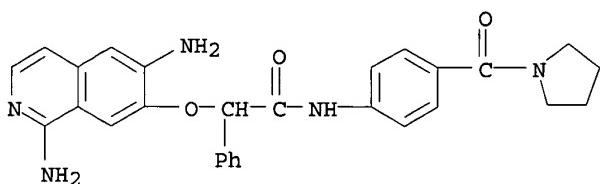
09/ 830,227

CN Benzeneacetamide, .alpha.-[(1-amino-6-hydroxy-7-isooquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)

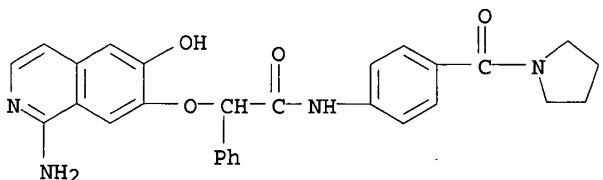


RN 489438-63-3 CAPLUS

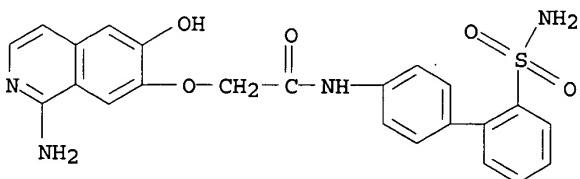
CN Benzeneacetamide, .alpha.-[(1,6-diamino-7-isooquinolinyl)oxy]-N-[4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



CN Benzeneacetamide, .alpha.-[(1-amino-6-hydroxy-7-isooquinolinyl)oxy]-N-[4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)

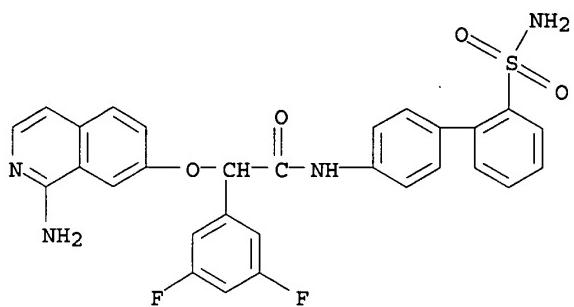


CN Acetamide, 2-[(1-amino-6-hydroxy-7-isooquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



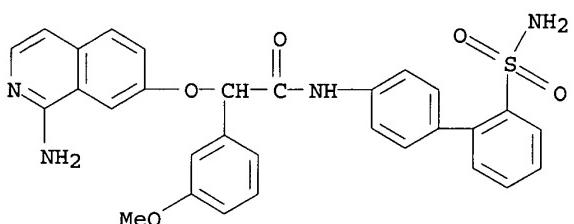
RN 489448-25-1 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isooquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]-3,5-difluoro- (9CI) (CA INDEX NAME)



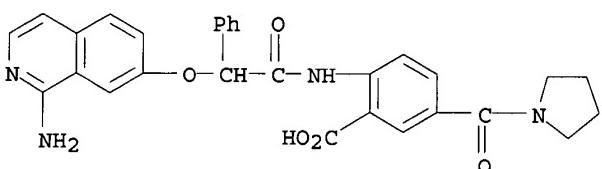
RN 489448-31-9 CAPLUS

CN Benzenacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]-3-methoxy- (9CI) (CA INDEX NAME)



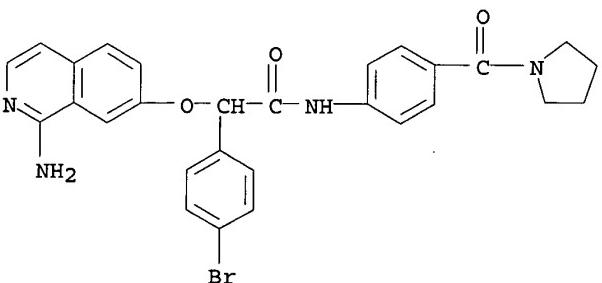
RN 489448-49-9 CAPLUS

CN Benzoic acid, 2-[[{(1-amino-7-isoquinolinyl)oxy}phenylacetyl]amino]-5-(1-pyrrolidinylcarbonyl)- (9CI) (CA INDEX NAME)



RN 489448-64-8 CAPLUS

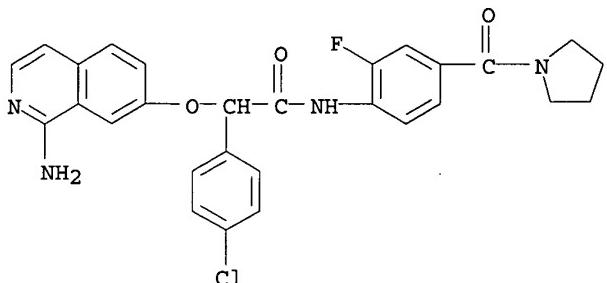
CN Benzenacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-4-bromo-N-[4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



09/ 830,227

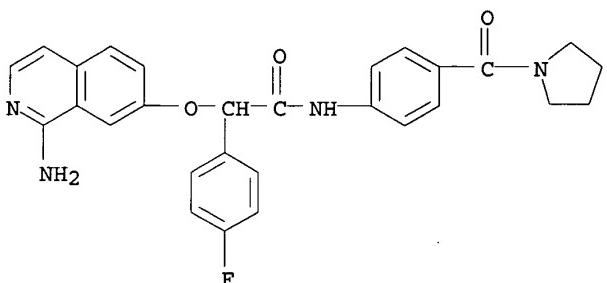
RN 489448-65-9 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-4-chloro-N-[2-fluoro-4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



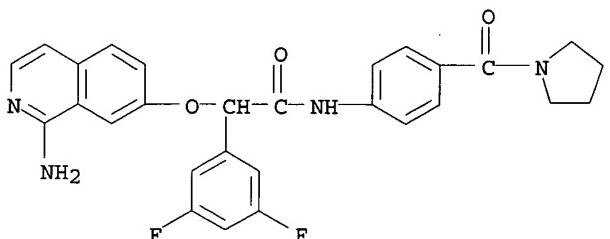
RN 489448-66-0 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-4-fluoro-N-[4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



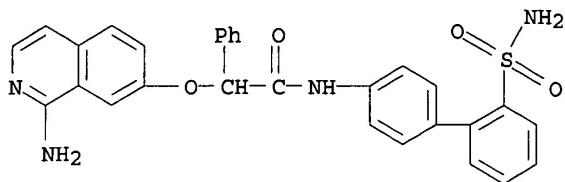
RN 489448-67-1 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-3,5-difluoro-N-[4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



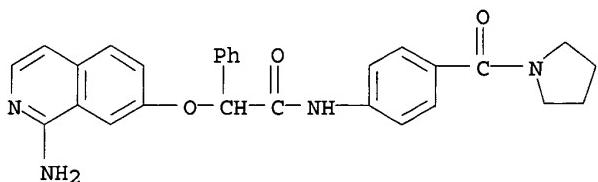
RN 308288-71-3 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



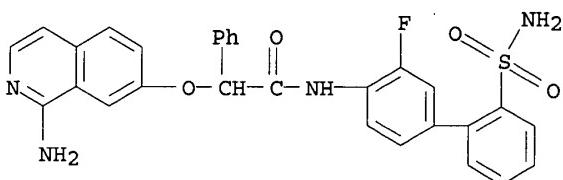
RN 308288-72-4 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



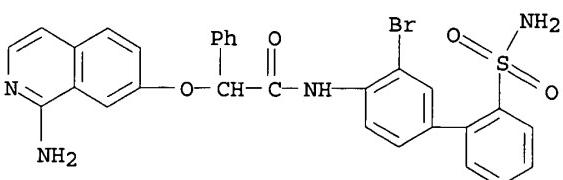
RN 308288-75-7 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)-3-fluoro[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



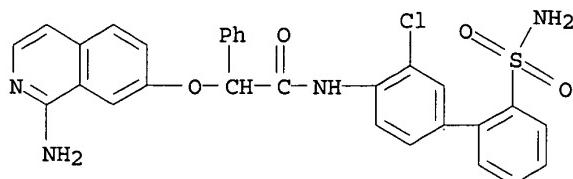
RN 308288-76-8 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)-3-bromo[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



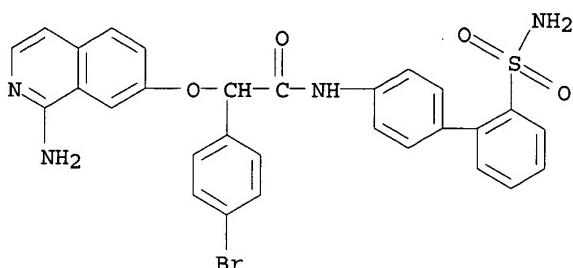
RN 308288-77-9 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)-3-chloro[1,1'-biphenyl]-4-yl]- (9CI) (CA INDEX NAME)



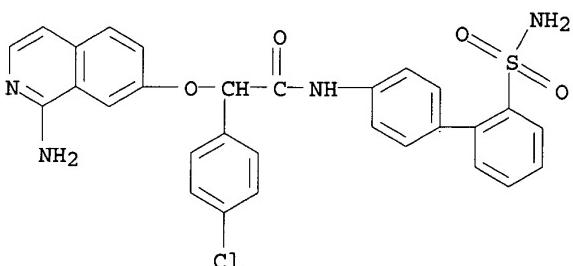
RN 308288-78-0 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]-4-bromo- (9CI) (CA INDEX NAME)



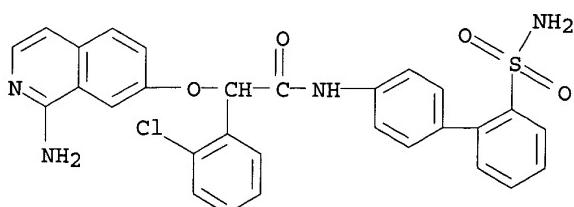
RN 308288-79-1 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]-4-chloro- (9CI) (CA INDEX NAME)



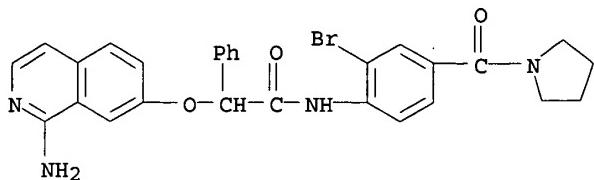
RN 308288-80-4 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]-2-chloro- (9CI) (CA INDEX NAME)



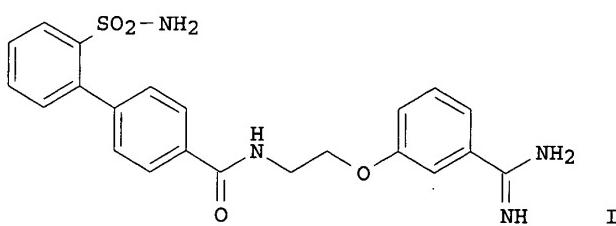
RN 308288-83-7 CAPLUS

CN Benzeneacetamide, .alpha.-[(1-amino-7-isoquinolinyl)oxy]-N-[2-bromo-4-(1-pyrrolidinylcarbonyl)phenyl]- (9CI) (CA INDEX NAME)



L3 ANSWER 8 OF 10 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 2000:842104 CAPLUS
 DOCUMENT NUMBER: 134:29204
 TITLE: Preparation of benzamidines and arylamidines as inhibitors of factor Xa
 INVENTOR(S): Zhu, Bing-Yan; Zhang, Penglie; Scarborough, Robert M.
 PATENT ASSIGNEE(S): Cor Therapeutics, Inc., USA
 SOURCE: PCT Int. Appl., 104 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 3
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|------------------|-----------------|------------|
| WO 2000071508 | A2 | 20001130 | WO 2000-US14208 | 20000524 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| EP 1185508 | A2 | 20020313 | EP 2000-932732 | 20000524 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| JP 2003500383 | T2 | 20030107 | JP 2000-619765 | 20000524 |
| PRIORITY APPLN. INFO.: | | | US 1999-135849P | P 19990524 |
| | | | WO 2000-US14208 | W 20000524 |
| OTHER SOURCE(S): GI | | MARPAT 134:29204 | | |



AB AYDEGJZL [wherein A = (cyclo)alkyl, (un)substituted amino, imino, amidino, guanidino, Ph, naphthyl, heterocyclic ring, etc.; Y = bond, CH₂, CO, NR₄CH₂, CH₂NR₄, NR₄, CONR₄, NR₄CO, C(:NR₄), C(:N₄)NR₄a, C(:NR₄)CH₂, C(:NR₄)NR₄ach₂, SO₂, O, SO₂NR₄, or NR₄SO₂; R₄ and R₄a = independently H,

alkyl, alkenyl, alkynyl, (alkyl)cycloalkyl, or (un)substituted alkylphenyl or alkynaphthyl; D = bond, (un)substituted Ph, naphthyl, or heterocyclic ring; E = NR₅CO, NR₅CONR₆, SO₂NR₅, NR₅SO₂NR₆, NR₅SO₂NR₆CO; R₅ and R₆ = H, alkyl, alkenyl, alkynyl, (alkyl)cycloalkyl or (un)substituted alkylphenyl, alkynaphthyl, alkylheteroaryl, carboxyalkyl, carbamidoalkyl, etc.; G = (un)substituted methylene, ethylene, or propylene; J = bond, CONR₁₁, NR₁₁CO, NR₁₁, NR₁₁CH₂, O, S, SO₂, SO, OCH₂, or SO₂CH₂; R₁₁ = H, alkyl, alkenyl, alkynyl, (alkyl)cycloalkyl or (un)substituted alkylphenyl, alkynaphthyl, or alkylheteroaryl; Z = (un)substituted Ph, naphthyl, or heterocyclic ring; L = H, CN, CONR₁₂NR₁₃, (CH₂)₀₋₂NR₁₂R₁₃, C(:NR₁₂)NR₁₂R₁₃, NR₁₂R₁₃, OR₁₂, NR₁₂C(:NR₁₂)NR₁₂N₃, or NR₁₂C(:N₁₂)R₁₃; R₁₂ and R₁₃ = independently H, OH, alkyl, (un)substituted alkoxy, (di)alkylamino, alkylphenyl, alkynaphthyl, carboxyalkyl, etc.] were prepd. as potent and highly selective inhibitors of factor Xa for the prevention or treatment of coagulation disorders (no data). For example, N-tert-butoxycarbonyl glycine was condensed with 3-cyanophenol in the presence of PPh₃ and DEAD in CH₂Cl₂ (93%), and the amine deprotected and converted to the salt using TFA. Reaction of the TFA amine salt with 2'-(tert-butylaminosulfonyl)-4-biphenylcarboxylic acid in the presence of BOP and i-Pr₂NET in DMF gave the amide (84%). The benzonitrile was converted to the desired benzamide salt (I.bul.TFA) in 85% yield by bubbling HCl gas through a soln. of the amide intermediate in MeOH, followed by neutralization and workup using 0.5% TFA in H₂O/MeCN. Compds. of the invention show selectivity for factor Xa vs. other proteases of the coagulation cascade or the fibrinolytic cascade, and are useful as diagnostic reagents as well as antithrombotic agents (no data).

IT 244256-82-4P 309930-02-7P 309930-03-8P

309930-04-9P 309930-05-0P 309930-06-1P

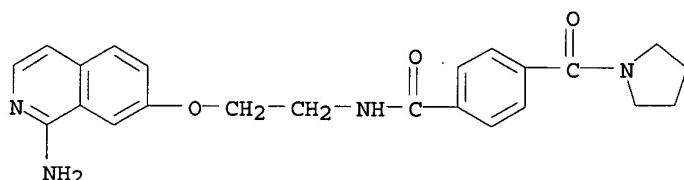
309930-07-2P 309930-09-4P 309930-30-1P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of benzamide and arylamide factor Xa inhibitors from benzonitriles and arylnitriles)

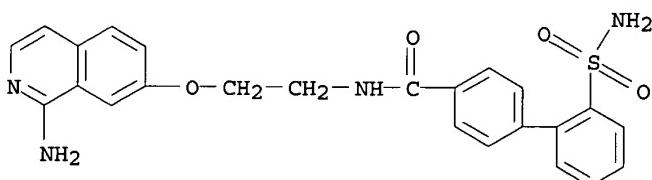
RN 244256-82-4 CAPLUS

CN Benzamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]ethyl]-4-(1-pyrrolidinylcarbonyl)- (9CI) (CA INDEX NAME)



RN 309930-02-7 CAPLUS

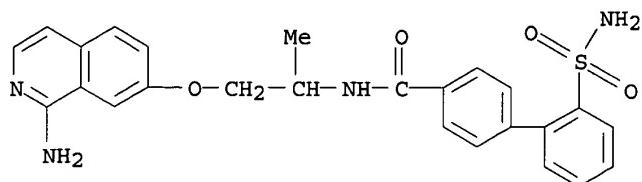
CN [1,1'-Biphenyl]-4-carboxamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]ethyl]-2'-(aminosulfonyl)- (9CI) (CA INDEX NAME)



RN 309930-03-8 CAPLUS

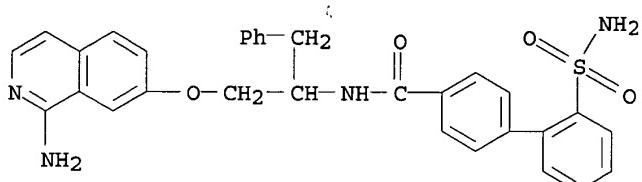
09/ 830,227

CN [1,1'-Biphenyl]-4-carboxamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]-1-methylethyl]-2'-(aminosulfonyl)- (9CI) (CA INDEX NAME)



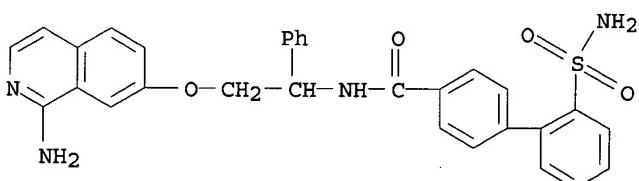
RN 309930-04-9 CAPLUS

CN [1,1'-Biphenyl]-4-carboxamide, N-[1-[(1-amino-7-isooquinolinyl)oxy]methyl]-2-phenylethyl]-2'-(aminosulfonyl)- (9CI) (CA INDEX NAME)



RN 309930-05-0 CAPLUS

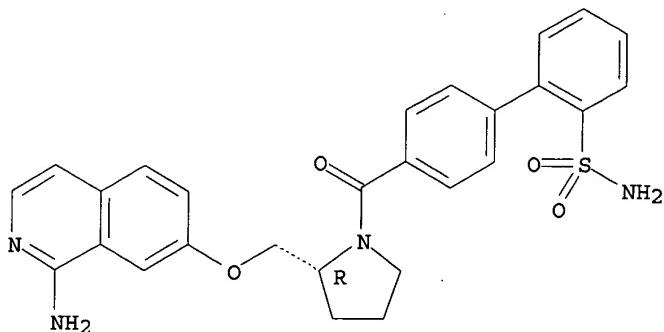
CN [1,1'-Biphenyl]-4-carboxamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]-1-phenylethyl]-2'-(aminosulfonyl)- (9CI) (CA INDEX NAME)



RN 309930-06-1 CAPLUS

CN Pyrrolidine, 2-[(1-amino-7-isooquinolinyl)oxy]methyl]-1-[[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]carbonyl]-, (2R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

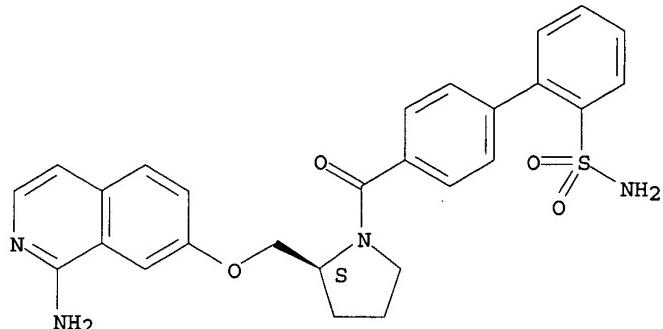


09/ 830,227

RN 309930-07-2 CAPLUS

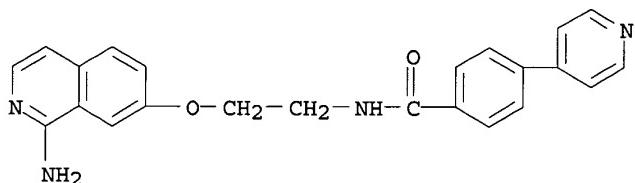
CN Pyrrolidine, 2-[(1-amino-7-isoquinolinyl)oxy]methyl]-1-[[2'-(aminosulfonyl)[1,1'-biphenyl]-4-yl]carbonyl]-, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



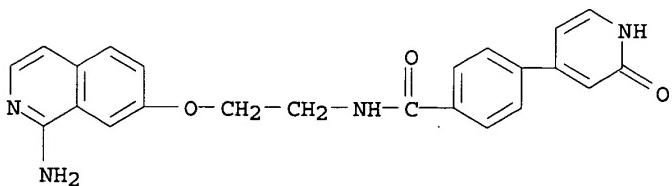
RN 309930-09-4 CAPLUS

CN Benzamide, N-[2-[(1-amino-7-isoquinolinyl)oxy]ethyl]-4-(4-pyridinyl)- (9CI) (CA INDEX NAME)



RN 309930-30-1 CAPLUS

CN Benzamide, N-[2-[(1-amino-7-isoquinolinyl)oxy]ethyl]-4-(1,2-dihydro-2-oxo-4-pyridinyl)- (9CI) (CA INDEX NAME)



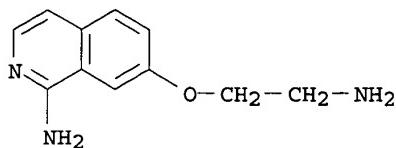
IT 309930-41-4P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(prepn. of benzamidine and arylamidine factor Xa inhibitors from benzonitriles and arylnitriles)

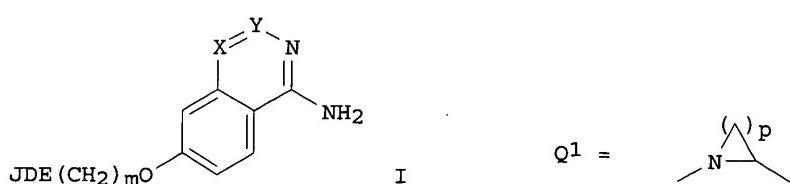
RN 309930-41-4 CAPLUS

CN 1-Isoquinolinamine, 7-(2-aminoethoxy)- (9CI) (CA INDEX NAME)



L3 ANSWER 9 OF 10 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 2000:291003 CAPLUS
 DOCUMENT NUMBER: 132:322143
 TITLE: Preparation of isoquinoline amino acid derivatives as serine protease inhibitors.
 INVENTOR(S): Timmers, Cornelis Marius; Rewinkel, Johannes Bernardus Maria
 PATENT ASSIGNEE(S): Akzo Nobel N.V., Neth.
 SOURCE: PCT Int. Appl., 38 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|------------|
| WO 2000024718 | A1 | 20000504 | WO 1999-EP7928 | 19991019 |
| W: AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GE, HU, ID, IL, IN, IS, JP, KP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 9963413 | A1 | 20000515 | AU 1999-63413 | 19991019 |
| BR 9914694 | A | 20010710 | BR 1999-14694 | 19991019 |
| EP 1123280 | A1 | 20010816 | EP 1999-950761 | 19991019 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| JP 2002528438 | T2 | 20020903 | JP 2000-578288 | 19991019 |
| NZ 511067 | A | 20030328 | NZ 1999-511067 | 19991019 |
| NO 2001001966 | A | 20010423 | NO 2001-1966 | 20010420 |
| PRIORITY APPLN. INFO.: | | | EP 1998-203559 | A 19981023 |
| | | | WO 1999-EP7928 | W 19991019 |
| OTHER SOURCE(S): MARPAT 132:322143 | | | | |
| GI | | | | |



AB Title compds. [I; J = H, R1, R1O2C, R1CO, R1SO2, etc.; D = NHCHR1CO, D-1-Tiq, D-Atc, Aic, D-1-Piq, etc.; E = NR2CH2, (substituted) Q1; R1 = (substituted) alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkylalkylene; R2 = H, R1; X, Y = CH, N, both may not = N; m = 1, 2; p = 2-4], were prepd.

Thus, (2S)-1-[N-(-)-camphorsulfonyl-D-cyclohexylalaninyl]-2-[2-(1-aminoisoquinolin-6-oxy)ethyl]piperidine (soln. phase prepn. given) showed antithrombin activity with IC₅₀ = 0.41. μ M.

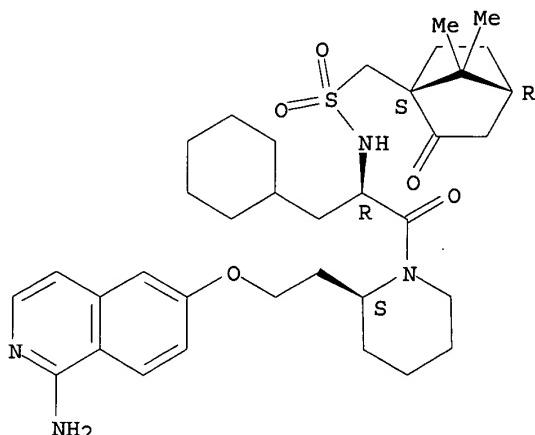
IT 266690-33-9P 266690-34-0P 266690-35-1P
 266690-36-2P 266690-37-3P 266690-38-4P
 266690-39-5P 266690-40-8P 266690-41-9P
 266690-42-0P 266690-43-1P 266690-44-2P
 266690-45-3P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (prepn. of isoquinoline amino acid derivs. as serine protease inhibitors)

RN 266690-33-9 CAPLUS

CN Piperidine, 2-[2-[(1-amino-6-isoquinolinyl)oxy]ethyl]-1-[(2R)-3-cyclohexyl-2-[[[(1S,4R)-7,7-dimethyl-2-oxobicyclo[2.2.1]hept-1-yl]methyl]sulfonyl]amino]-1-oxopropyl]-, (2S)- (9CI) (CA INDEX NAME)

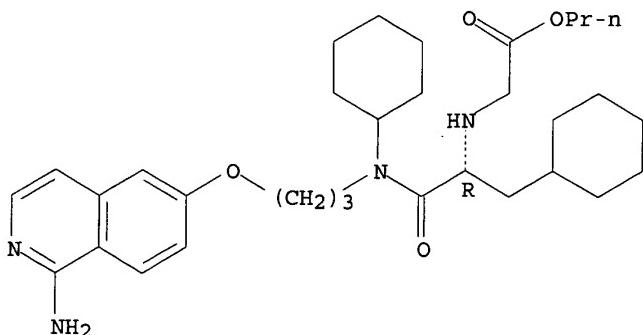
Absolute stereochemistry.



RN 266690-34-0 CAPLUS

CN Glycine, N-[(1R)-2-[[3-[(1-amino-6-isoquinolinyl)oxy]propyl]cyclohexylamino]-1-(cyclohexylmethyl)-2-oxoethyl]-, propyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

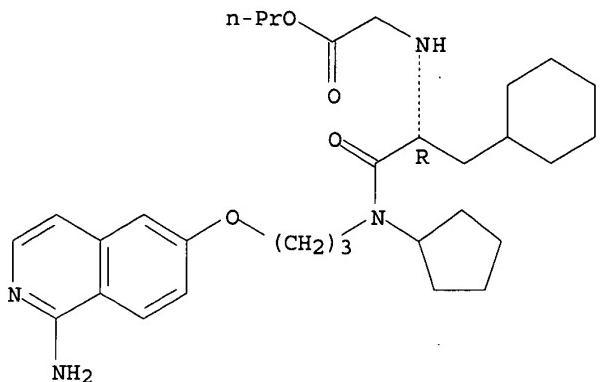


RN 266690-35-1 CAPLUS

CN Glycine, N-[(1R)-2-[[3-[(1-amino-6-isoquinolinyl)oxy]propyl]cyclopentylamino]-1-(cyclohexylmethyl)-2-oxoethyl]-, propyl ester (9CI) (CA INDEX NAME)

09/ 830,227

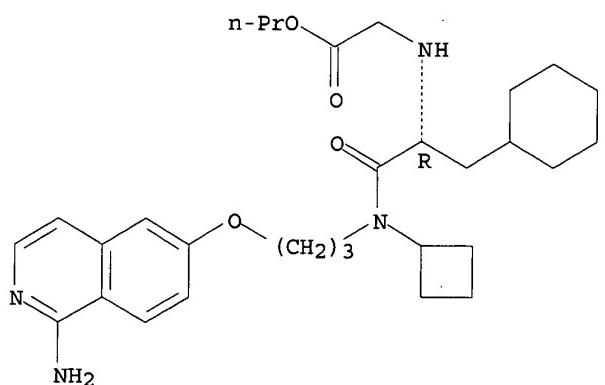
Absolute stereochemistry.



RN 266690-36-2 CAPLUS

CN Glycine, N-[(1R)-2-[[3-[(1-amino-6-isoquinolinyl)oxy]propyl]cyclobutylamino]-1-(cyclohexylmethyl)-2-oxoethyl]-, propyl ester (9CI) (CA INDEX NAME)

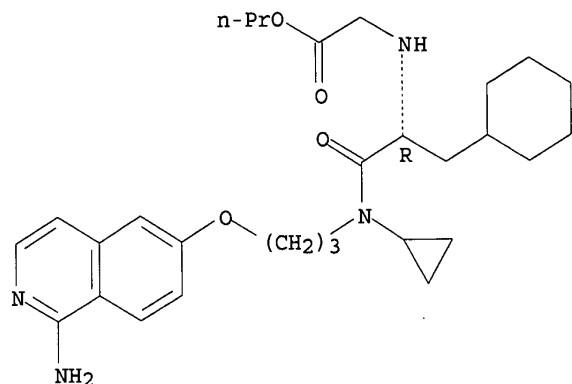
Absolute stereochemistry.



RN 266690-37-3 CAPLUS

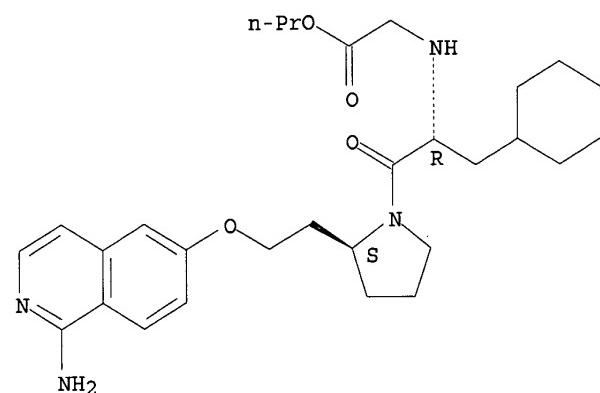
CN Glycine, N-[(1R)-2-[[3-[(1-amino-6-isoquinolinyl)oxy]propyl]cyclopropylamino]-1-(cyclohexylmethyl)-2-oxoethyl]-, propyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.



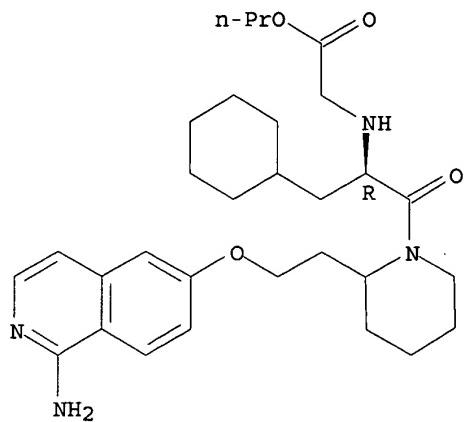
RN 266690-38-4 CAPLUS
CN Glycine, N-[(1R)-2-[(2S)-2-[2-[(1-amino-6-isoquinolinyl)oxy]ethyl]-1-pyrrolidinyl]-1-(cyclohexylmethyl)-2-oxoethyl]-, propyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 266690-39-5 CAPLUS
CN Glycine, N-[(1R)-2-[(2-[(1-amino-6-isoquinolinyl)oxy]ethyl)-1-piperidinyl]-1-(cyclohexylmethyl)-2-oxoethyl]-, propyl ester (9CI) (CA INDEX NAME)

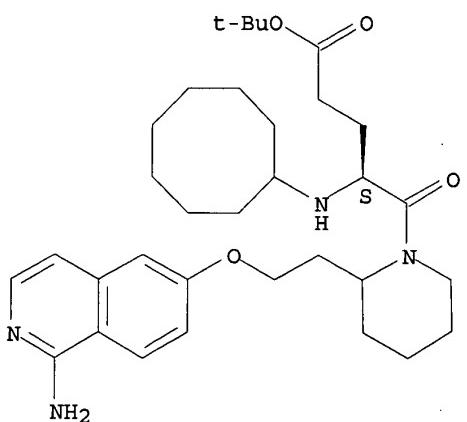
Absolute stereochemistry.



RN 266690-40-8 CAPLUS

CN 1-Piperidinpentanoic acid, 2-[2-[(1-amino-6-isoquinolinyloxy)ethyl]-.gamma.- (cyclooctylamino)-.delta.-oxo-, 1,1-dimethylethyl ester, (.gamma.S)- (9CI) (CA INDEX NAME)

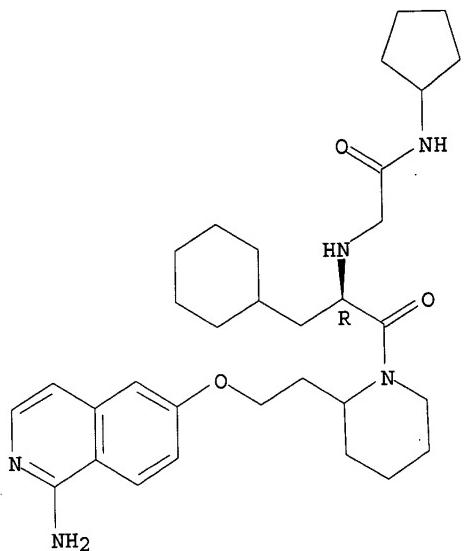
Absolute stereochemistry.



RN 266690-41-9 CAPLUS

CN Acetamide, 2-[[1R]-2-[2-[(1-amino-6-isoquinolinyloxy)ethyl]-1-piperidinyl]-1-(cyclohexylmethyl)-2-oxoethyl]amino]-N-cyclopentyl- (9CI) (CA INDEX NAME)

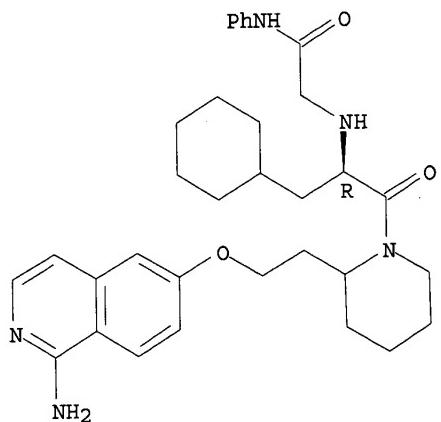
Absolute stereochemistry.



RN 266690-42-0 CAPLUS

CN Acetamide, 2-[(1R)-2-[2-[(1-amino-6-isoquinolinyl)oxy]ethyl]-1-piperidinyl]-1-(cyclohexylmethyl)-2-oxoethylamino]-N-phenyl- (9CI) (CA INDEX NAME)

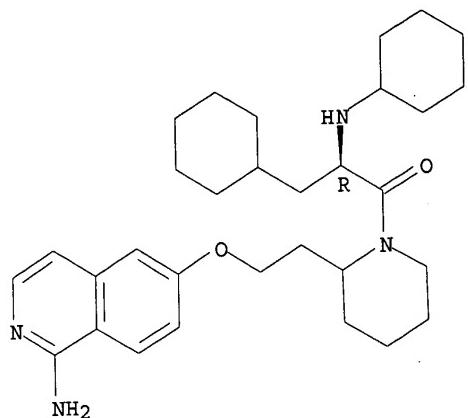
Absolute stereochemistry.



RN 266690-43-1 CAPLUS

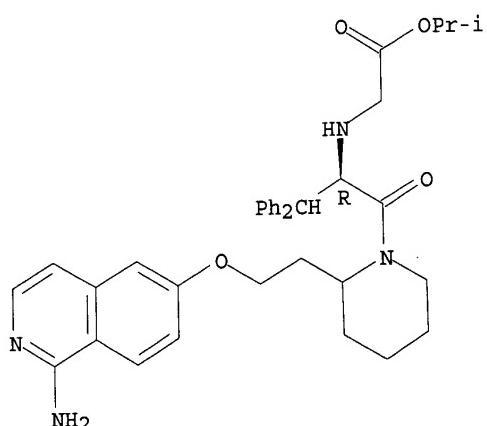
CN Piperidine, 2-[2-[(1-amino-6-isoquinolinyl)oxy]ethyl]-1-[(2R)-3-cyclohexyl-2-(cyclohexylamino)-1-oxopropyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



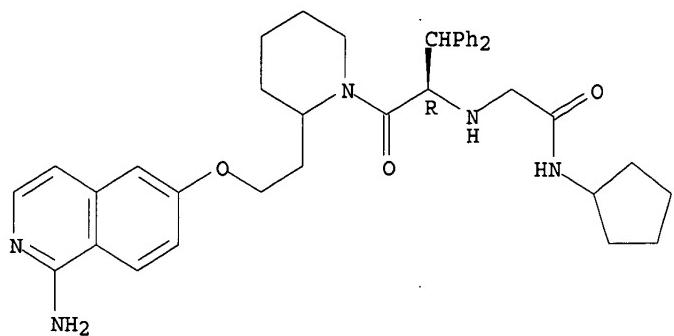
RN 266690-44-2 CAPLUS
CN Glycine, N-[(1R)-1-[[2-[2-[(1-amino-6-isoquinolinyl)oxy]ethyl]-1-piperidinyl]carbonyl]-2,2-diphenylethyl]-, 1-methylethyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 266690-45-3 CAPLUS
CN Acetamide, 2-[(1R)-1-[[2-[2-[(1-amino-6-isoquinolinyl)oxy]ethyl]-1-piperidinyl]carbonyl]-2,2-diphenylethyl]amino]-N-cyclopentyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

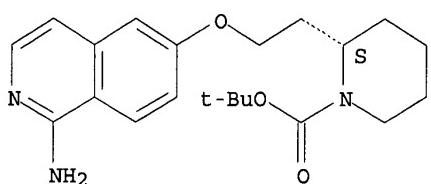


IT 266690-50-0P 266690-56-6P 266690-64-6P
 266690-69-1P 266690-72-6P 266690-75-9P
 266690-88-4P 266690-89-5P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
 (Reactant or reagent)
 (prepn. of isoquinoline amino acid derivs. as serine protease
 inhibitors)

RN 266690-50-0 CAPPLUS

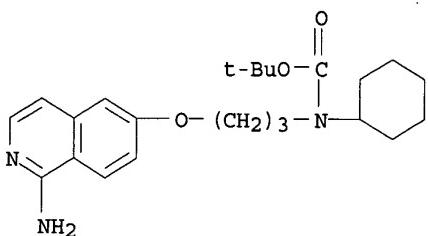
CN 1-Piperidinecarboxylic acid, 2-[2-[(1-amino-6-isoquinolinyl)oxy]ethyl]-,
 1,1-dimethylethyl ester, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



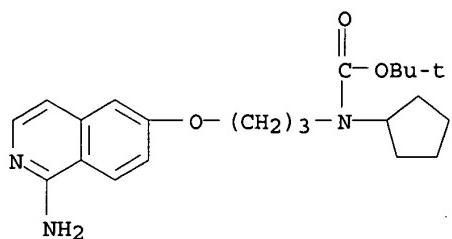
RN 266690-56-6 CAPPLUS

CN Carbamic acid, [3-[(1-amino-6-isoquinolinyl)oxy]propyl]cyclohexyl-,
 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



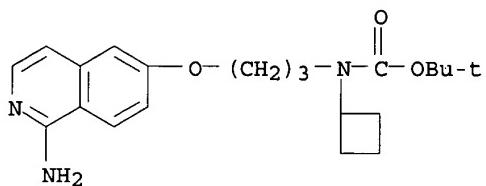
RN 266690-64-6 CAPPLUS

CN Carbamic acid, [3-[(1-amino-6-isoquinolinyl)oxy]propyl]cyclopentyl-,
 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



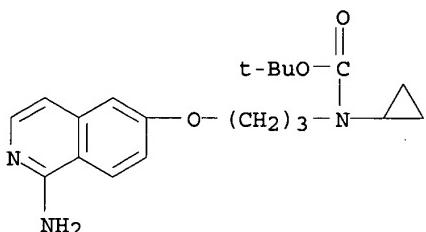
RN 266690-69-1 CAPLUS

CN Carbamic acid, [3-[(1-amino-6-isoquinolinyl)oxy]propyl]cyclobutyl-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



RN 266690-72-6 CAPLUS

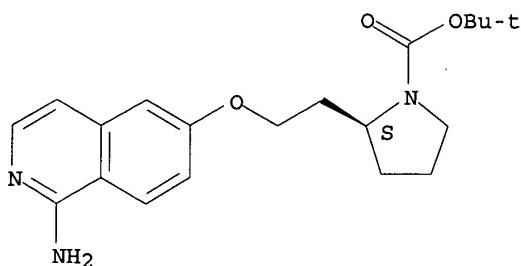
CN Carbamic acid, [3-[(1-amino-6-isoquinolinyl)oxy]propyl]cyclopropyl-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



RN 266690-75-9 CAPLUS

CN 1-Pyrrolidinecarboxylic acid, 2-[2-[(1-amino-6-isoquinolinyl)oxy]ethyl]-, 1,1-dimethylethyl ester, (2S)- (9CI) (CA INDEX NAME)

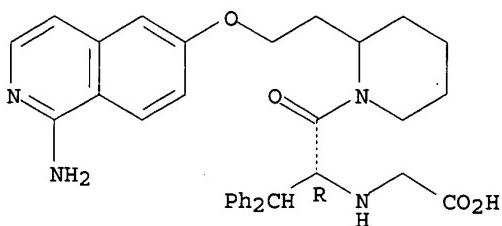
Absolute stereochemistry.



RN 266690-88-4 CAPLUS

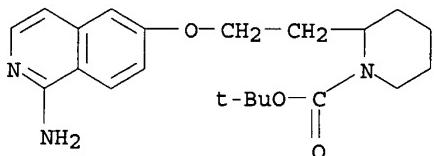
CN Glycine, N-[(1R)-1-[[2-[(1-amino-6-isoquinolinyl)oxy]ethyl]-1-piperidinyl]carbonyl]-2,2-diphenylethyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 266690-89-5 CAPLUS

CN 1-Piperidinecarboxylic acid, 2-[2-[(1-amino-6-isoquinolinyl)oxy]ethyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 10 OF 10 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 1999:613871 CAPLUS
 DOCUMENT NUMBER: 131:243189
 TITLE: Preparation of aminoisoquinoline derivatives as inhibitors of activated blood coagulation factor X
 INVENTOR(S): Nakagawa, Tadakiyo; Makino, Shingo; Sagi, Kazuyuki; Takayanagi, Masaru; Kayahara, Takashi; Takehana, Shunji
 PATENT ASSIGNEE(S): Ajinomoto Co., Inc., Japan
 SOURCE: PCT Int. Appl., 80 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 9947503 | A1 | 19990923 | WO 1999-JP1309 | 19990317 |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2324153 | AA | 19990923 | CA 1999-2324153 | 19990317 |
| AU 9928522 | A1 | 19991011 | AU 1999-28522 | 19990317 |
| AU 753675 | B2 | 20021024 | | |
| EP 1065200 | A1 | 20010103 | EP 1999-909191 | 19990317 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, NL, SE, PT, IE, FI | | | | |

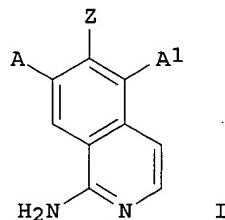
09/ 830,227

PRIORITY APPLN. INFO.:

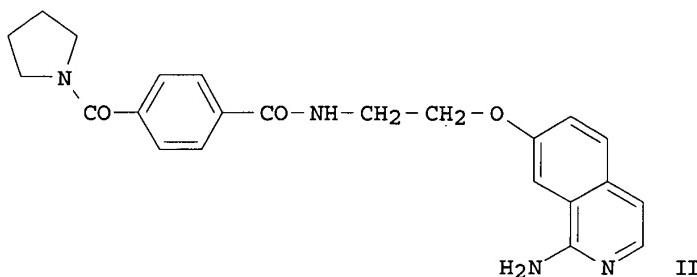
JP 1998-70771 A 19980319
JP 1998-197133 A 19980713
WO 1999-JP1309 W 19990317

OTHER SOURCE(S):
GI

MARPAT 131:243189



I



II

AB The title compds. I [A is VLY, A1 is H; or A1 is VLY, A is H ; L is CH₂CH₂, etc.; V is, for example, H, (un)substituted benzoyl, etc.; extensive details on V are given; Y is CH:CH, etc.; Z = H, alkyl, etc.] are prep'd. I are useful as active ingredients in anticoagulants or preventives/remedies for thrombosis or embolism. In an in vitro test for inhibition of the activated blood coagulation factor X, the title compd. II showed pIC₅₀ of 6.6.

IT 244256-81-3P 244256-83-5P 244256-85-7P
244256-87-9P 244256-89-1P 244256-91-5P
244256-93-7P 244256-95-9P 244256-97-1P
244256-99-3P 244257-01-0P 244257-03-2P
244257-05-4P 244257-07-6P 244257-09-8P
244257-11-2P 244257-13-4P 244257-15-6P
244257-17-8P 244257-19-0P 244257-21-4P
244257-23-6P 244257-25-8P 244257-27-0P
244257-29-2P 244257-31-6P 244257-33-8P
244257-35-0P 244257-37-2P 244257-39-4P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(prep'n. of aminoisoquinoline derivs. as inhibitors of activated blood coagulation factor X)

RN 244256-81-3 CAPLUS

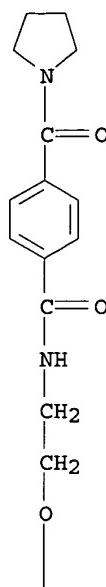
CN Benzamide, N-[2-[(1-amino-5-isoquinolinyl)oxy]ethyl]-4-(1-pyrrolidinylcarbonyl)-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

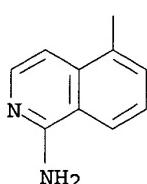
CRN 244256-80-2

CMF C23 H24 N4 O3

PAGE 1-A

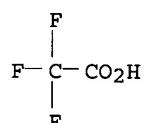


PAGE 2-A



CM 2

CRN 76-05-1
CMF C₂ H F₃ O₂



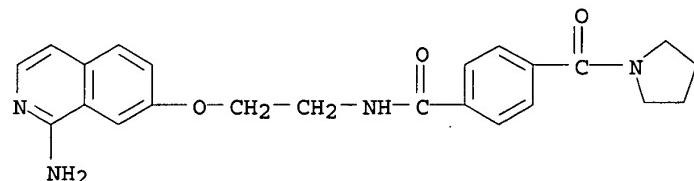
RN 244256-83-5 CAPLUS
CN Benzamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]ethyl]-4-(1-pyrrolidinylcarbonyl)-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244256-82-4

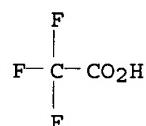
09/ 830,227

CMF C23 H24 N4 O3



CM 2

CRN 76-05-1
CMF C2 H F3 O2

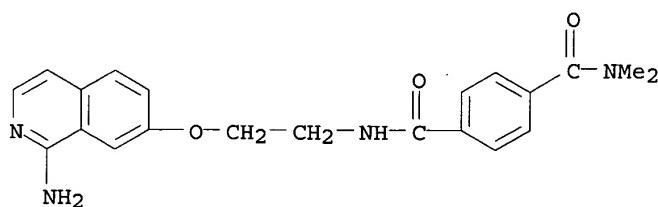


RN 244256-85-7 CAPLUS

CN 1,4-Benzenedicarboxamide, N'-[2-[(1-amino-7-isooquinolinyl)oxy]ethyl]-N,N-dimethyl-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

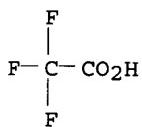
CM 1

CRN 244256-84-6
CMF C21 H22 N4 O3



CM 2

CRN 76-05-1
CMF C2 H F3 O2



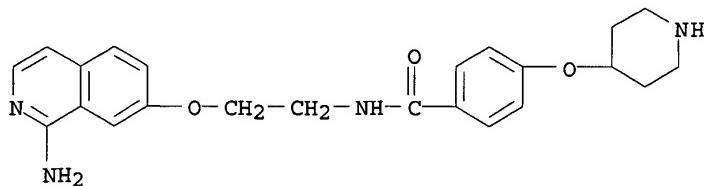
RN 244256-87-9 CAPLUS

CN Benzamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]ethyl]-4-(4-piperidinyloxy)-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

09/ 830,227

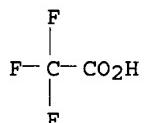
CM 1

CRN 244256-86-8
CMF C23 H26 N4 O3



CM 2

CRN 76-05-1
CMF C2 H F3 O2

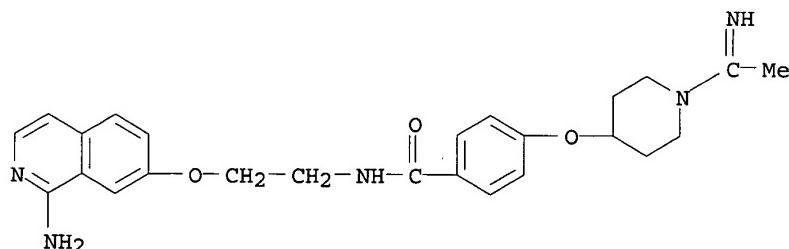


RN 244256-89-1 CAPLUS

CN Benzamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]ethyl]-4-[[1-(1-iminoethyl)-4-piperidinyl]oxy]-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

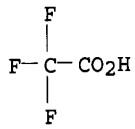
CM 1

CRN 244256-88-0
CMF C25 H29 N5 O3



CM 2

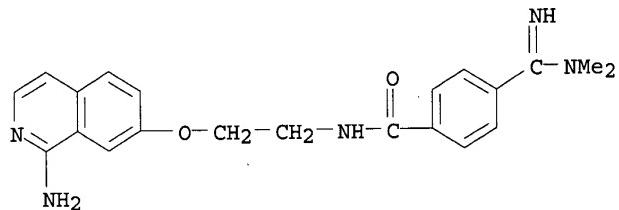
CRN 76-05-1
CMF C2 H F3 O2



RN 244256-91-5 CAPLUS
 CN Benzamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]ethyl]-4-[(dimethylamino)iminomethyl]-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

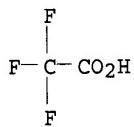
CM 1

CRN 244256-90-4
 CMF C21 H23 N5 O2



CM 2

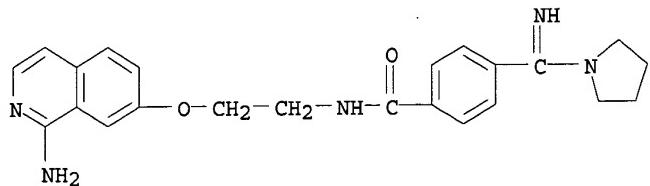
CRN 76-05-1
 CMF C2 H F3 O2



RN 244256-93-7 CAPLUS
 CN Benzamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]ethyl]-4-(imino-1-pyrrolidinylmethyl)-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

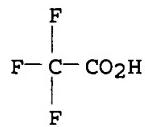
CRN 244256-92-6
 CMF C23 H25 N5 O2



09/ 830,227

CM 2

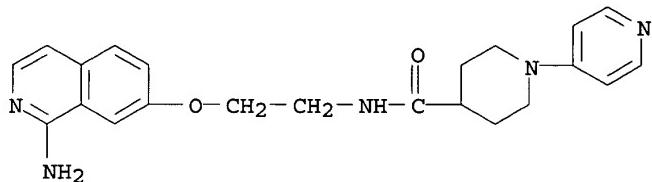
CRN 76-05-1
CMF C2 H F3 O2



RN 244256-95-9 CAPLUS
CN 4-Piperidinecarboxamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]ethyl]-1-(4-pyridinyl)-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

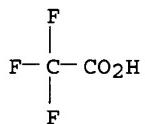
CM 1

CRN 244256-94-8
CMF C22 H25 N5 O2



CM 2

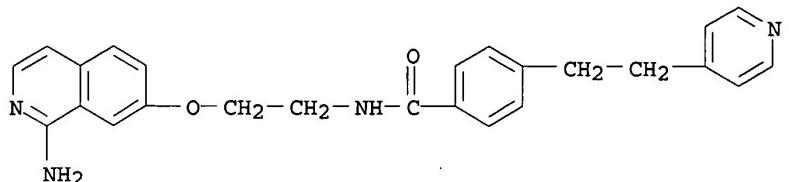
CRN 76-05-1
CMF C2 H F3 O2



RN 244256-97-1 CAPLUS
CN Benzamide, N-[2-[(1-amino-7-isooquinolinyl)oxy]ethyl]-4-[2-(4-pyridinyl)ethyl]-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

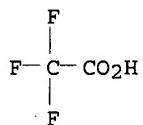
CM 1

CRN 244256-96-0
CMF C25 H24 N4 O2



CM 2

CRN 76-05-1
CMF C2 H F3 O2

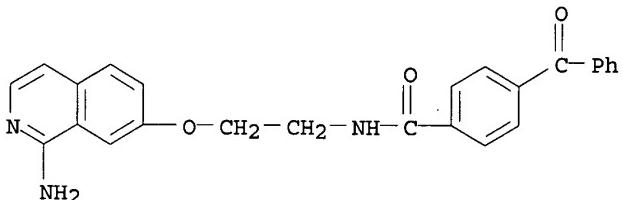


RN 244256-99-3 CAPLUS

CN Benzamide, N-[2-[(1-amino-7-isoquinolinyl)oxy]ethyl]-4-benzoyl-,
mono(trifluoroacetate) (9CI) (CA INDEX NAME)

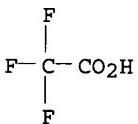
CM 1

CRN 244256-98-2
CMF C25 H21 N3 O3



CM 2

CRN 76-05-1
CMF C2 H F3 O2



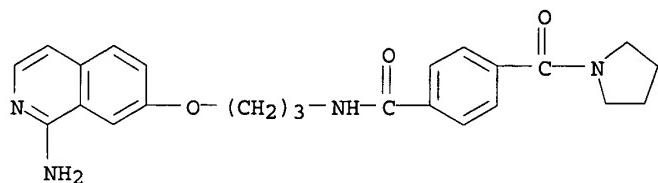
RN 244257-01-0 CAPLUS

CN Benzamide, N-[3-[(1-amino-7-isooquinolinyl)oxy]propyl]-4-(1-pyrrolidinylcarbonyl)-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

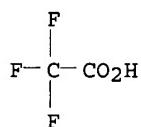
09/ 830,227

CRN 244257-00-9
CMF C24 H26 N4 O3



CM 2

CRN 76-05-1
CMF C2 H F3 O2

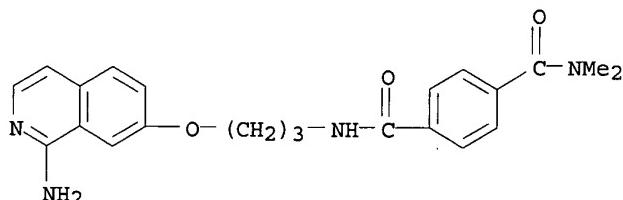


RN 244257-03-2 CAPLUS

CN 1,4-Benzenedicarboxamide, N'-(3-[(1-amino-7-isoquinolinyl)oxy]propyl)-N,N-dimethyl-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

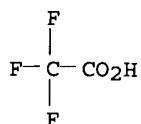
CM 1

CRN 244257-02-1
CMF C22 H24 N4 O3



CM 2

CRN 76-05-1
CMF C2 H F3 O2



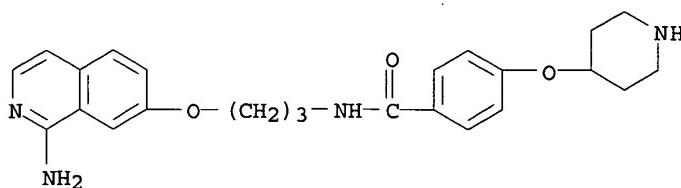
RN 244257-05-4 CAPLUS

09/ 830,227

CN Benzamide, N-[3-[(1-amino-7-isoquinolinyl)oxy]propyl]-4-(4-piperidinyloxy)-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

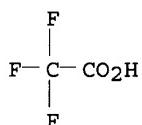
CM 1

CRN 244257-04-3
CMF C24 H28 N4 O3



CM 2

CRN 76-05-1
CMF C2 H F3 O2

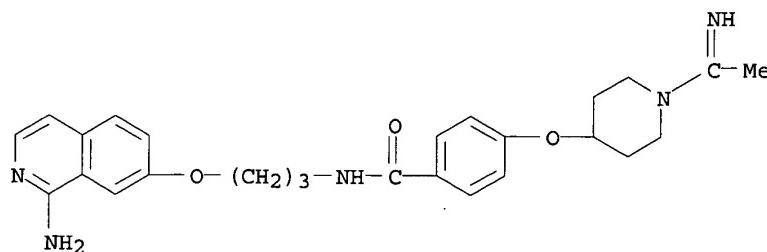


RN 244257-07-6 CAPLUS

CN Benzamide, N-[3-[(1-amino-7-isoquinolinyl)oxy]propyl]-4-[[1-(1-iminoethyl)-4-piperidinyloxy]-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

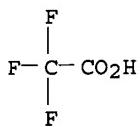
CM 1

CRN 244257-06-5
CMF C26 H31 N5 O3



CM 2

CRN 76-05-1
CMF C2 H F3 O2



RN 244257-09-8 CAPLUS

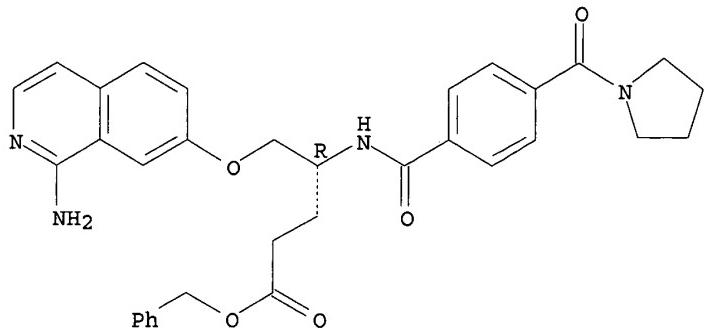
CN Pentanoic acid, 5-[(1-amino-7-isoquinolinyl)oxy]-4-[[4-(1-pyrrolidinylcarbonyl)benzoyl]amino]-, phenylmethyl ester, (4R)-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-08-7

CMF C33 H34 N4 O5

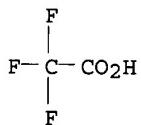
Absolute stereochemistry.



CM 2

CRN 76-05-1

CMF C2 H F3 O2



RN 244257-11-2 CAPLUS

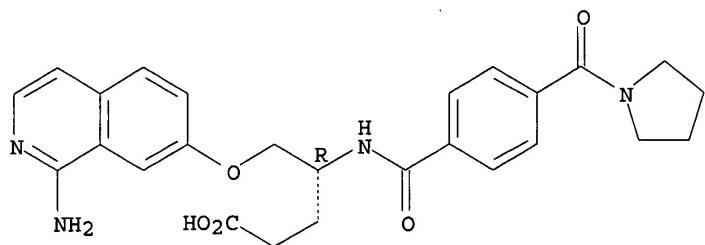
CN Pentanoic acid, 5-[(1-amino-7-isoquinolinyl)oxy]-4-[[4-(1-pyrrolidinylcarbonyl)benzoyl]amino]-, (4R)-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-10-1

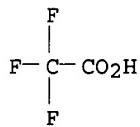
CMF C26 H28 N4 O5

Absolute stereochemistry.



CM 2

CRN 76-05-1
CMF C2 H F3 O2



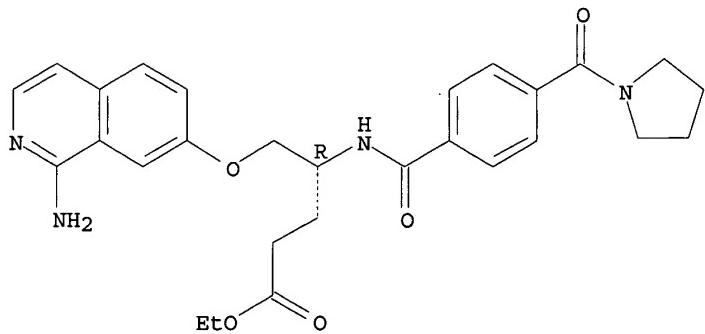
RN 244257-13-4 CAPLUS

CN Pentanoic acid, 5-[(1-amino-7-isoquinolinyl)oxy]-4-[[4-(1-pyrrolidinylcarbonyl)benzoyl]amino]-, ethyl ester, (4R)-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-12-3
CMF C28 H32 N4 O5

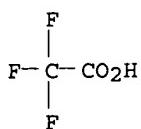
Absolute stereochemistry.



CM 2

CRN 76-05-1
CMF C2 H F3 O2

09/ 830,227



RN 244257-15-6 CAPLUS

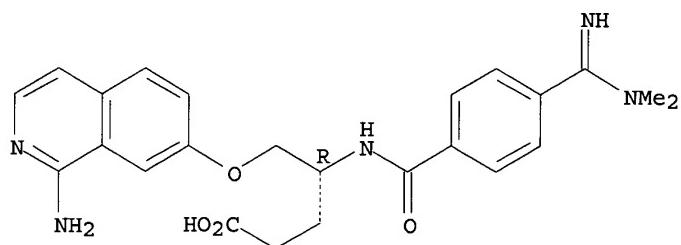
CN Pentanoic acid, 5-[(1-amino-7-isoquinolinyl)oxy]-4-[[4-[(dimethylamino)iminomethyl]benzoyl]amino]-, (4R)-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-14-5

CMF C24 H27 N5 O4

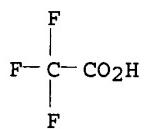
Absolute stereochemistry.



CM 2

CRN 76-05-1

CMF C2 H F3 O2



RN 244257-17-8 CAPLUS

CN Pentanoic acid, 5-[(1-amino-7-isoquinolinyl)oxy]-4-[[4-[(2-(4-pyridinyl)ethyl]benzoyl]amino]-, (4R)-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

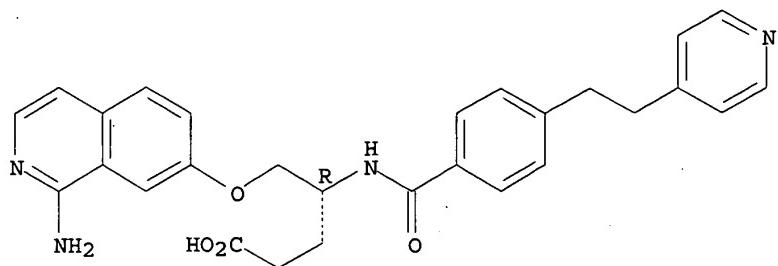
CM 1

CRN 244257-16-7

CMF C28 H28 N4 O4

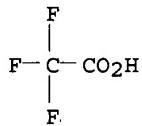
Absolute stereochemistry.

09/ 830,227



CM 2

CRN 76-05-1
CMF C2 H F3 O2

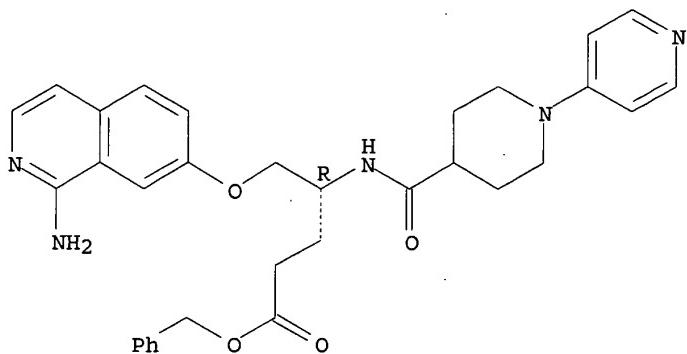


RN 244257-19-0 CAPLUS
CN Pentanoic acid, 5-[(1-amino-7-isoquinolinyl)oxy]-4-[[[1-(4-pyridinyl)-4-piperidinyl]carbonyl]amino]-, phenylmethyl ester, (4R)-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

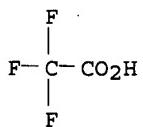
CRN 244257-18-9
CMF C32 H35 N5 O4

Absolute stereochemistry.



CM 2

CRN 76-05-1
CMF C2 H F3 O2



RN 244257-21-4 CAPLUS

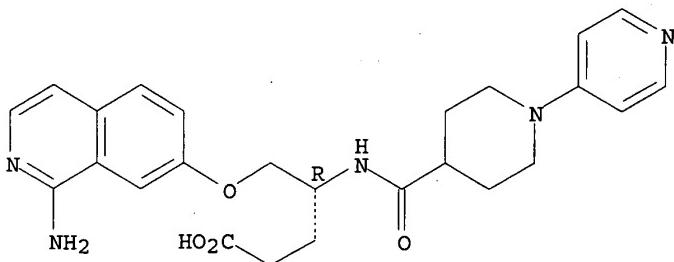
CN Pentanoic acid, 5-[(1-amino-7-isoquinolinyl)oxy]-4-[[[1-(4-pyridinyl)-4-piperidinyl]carbonyl]amino]-, (4R)-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-20-3

CMF C25 H29 N5 O4

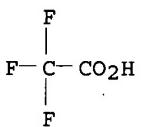
Absolute stereochemistry.



CM 2

CRN 76-05-1

CMF C2 H F3 O2



RN 244257-23-6 CAPLUS

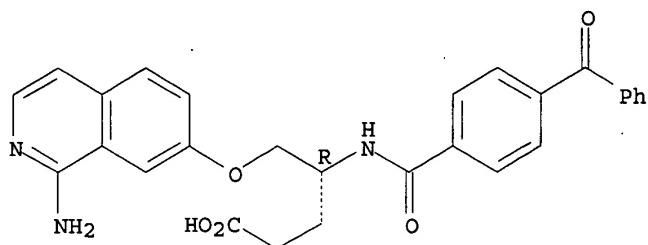
CN Pentanoic acid, 5-[(1-amino-7-isoquinolinyl)oxy]-4-[(4-benzoylbenzoyl)amino]-, (4R)-, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-22-5

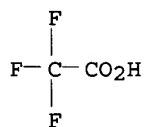
CMF C28 H25 N3 O5

Absolute stereochemistry.



CM 2

CRN 76-05-1
CMF C2 H F3 O2

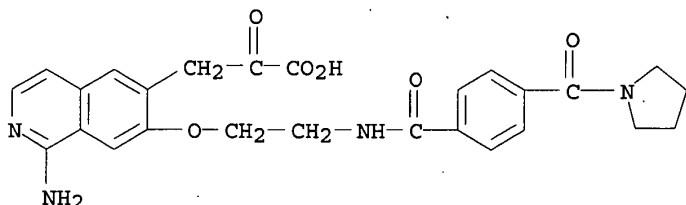


RN 244257-25-8 CAPLUS

CN 6-Isoquinolinepropanoic acid, 1-amino-.alpha.-oxo-7-[2-[[4-(1-pyrrolidinylcarbonyl)benzoyl]amino]ethoxy]-, mono(trifluoroacetate) (9CI)
(CA INDEX NAME)

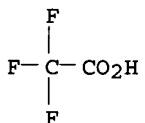
CM 1

CRN 244257-24-7
CMF C26 H26 N4 O6

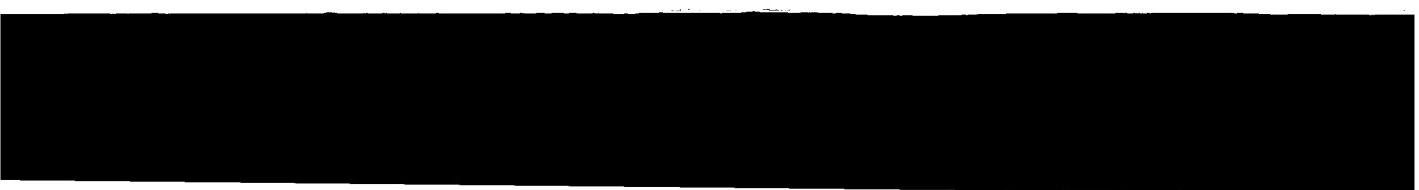


CM 2

CRN 76-05-1
CMF C2 H F3 O2



RN 244257-27-0 CAPLUS

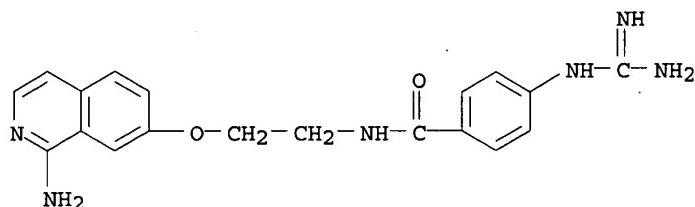


09/ 830,227

CN Benzamide, 4-[(aminoiminomethyl)amino]-N-[2-[(1-amino-7-isoquinolinyl)oxy]ethyl]-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

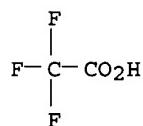
CM 1

CRN 244257-26-9
CMF C19 H20 N6 O2



CM 2

CRN 76-05-1
CMF C2 H F3 O2



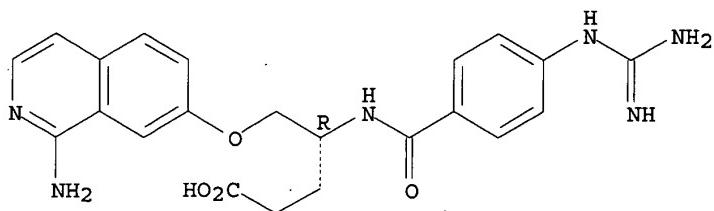
RN 244257-29-2 CAPLUS

CN Pentanoic acid, 4-[[4-[(aminoiminomethyl)amino]benzoyl]amino]-5-[(1-amino-7-isoquinolinyl)oxy]-, (4R)-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-28-1
CMF C22 H24 N6 O4

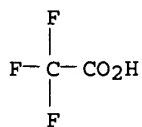
Absolute stereochemistry.



CM 2

CRN 76-05-1
CMF C2 H F3 O2

09/ 830,227



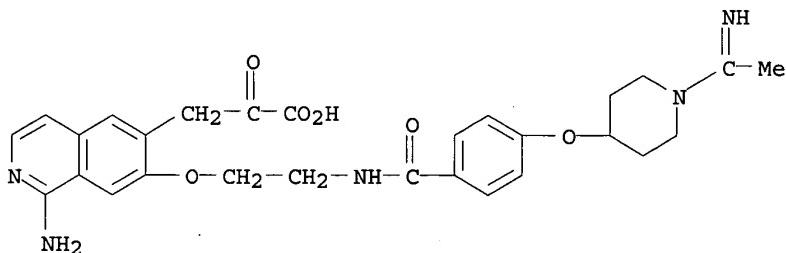
RN 244257-31-6 CAPLUS

CN 6-Isoquinolinepropanoic acid, 1-amino-7-[2-[[4-[[1-(1-iminoethyl)-4-piperidinyl]oxy]benzoyl]amino]ethoxy]-.alpha.-oxo-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-30-5

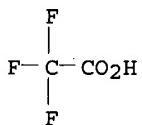
CMF C28 H31 N5 O6



CM 2

CRN 76-05-1

CMF C2 H F3 O2



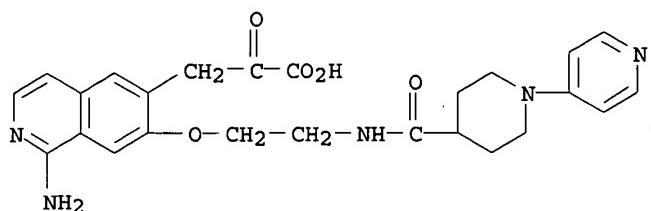
RN 244257-33-8 CAPLUS

CN 6-Isoquinolinepropanoic acid, 1-amino-.alpha.-oxo-7-[2-[[[1-(4-pyridinyl)-4-piperidinyl]carbonyl]amino]ethoxy]-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

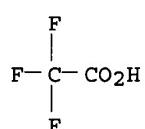
CRN 244257-32-7

CMF C25 H27 N5 O5



CM 2

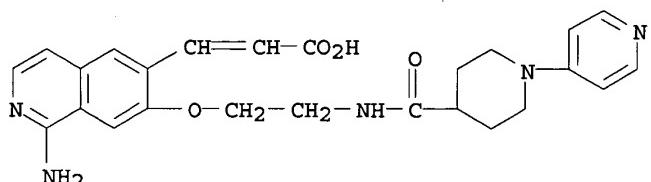
CRN 76-05-1
CMF C2 H F3 O2



RN 244257-35-0 CAPLUS
CN 2-Propenoic acid, 3-[1-amino-7-[2-[[1-(4-pyridinyl)-4-piperidinyl]carbonyl]amino]ethoxy]-6-isooquinolinyl-,
bis(trifluoroacetate) (9CI) (CA INDEX NAME)

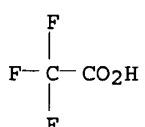
CM 1

CRN 244257-34-9
CMF C25 H27 N5 O4



CM 2

CRN 76-05-1
CMF C2 H F3 O2

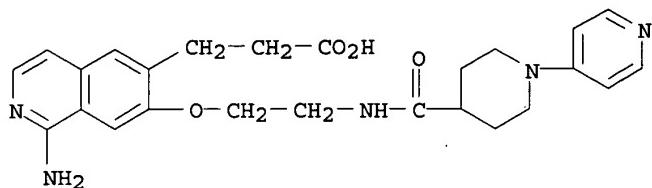


RN 244257-37-2 CAPLUS
CN 6-Isoquinolinepropanoic acid, 1-amino-7-[2-[[1-(4-pyridinyl)-4-piperidinyl]carbonyl]amino]ethoxy-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

09/ 830,227

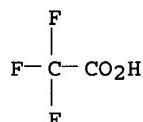
CM 1

CRN 244257-36-1
CMF C25 H29 N5 O4



CM 2

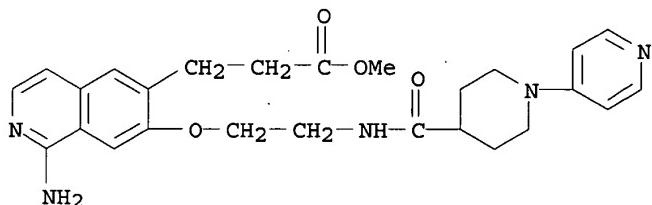
CRN 76-05-1
CMF C2 H F3 O2



RN 244257-39-4 CAPLUS
CN 6-Isoquinolinepropanoic acid, 1-amino-7-[2-[[[1-(4-pyridinyl)-4-piperidinyl]carbonyl]amino]ethoxy]-, methyl ester, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

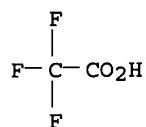
CM 1

CRN 244257-38-3
CMF C26 H31 N5 O4



CM 2

CRN 76-05-1
CMF C2 H F3 O2



IT 244257-45-2P 244257-53-2P 244257-58-7P
 244257-60-1P 244257-66-7P 244257-68-9P
 244257-70-3P 244257-72-5P 244257-74-7P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (prepn. of aminoisoquinoline derivs. as inhibitors of activated blood coagulation factor X)

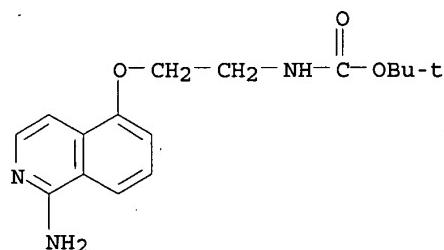
RN 244257-45-2 CAPLUS

CN Carbamic acid, [2-[(1-amino-5-isoquinolinyl)oxy]ethyl]-, 1,1-dimethylethyl ester, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-44-1

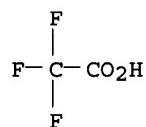
CMF C16 H21 N3 O3



CM 2

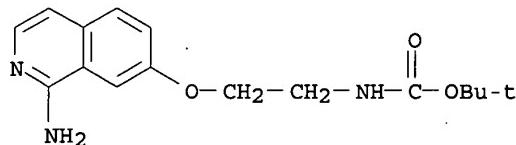
CRN 76-05-1

CMF C2 H F3 O2



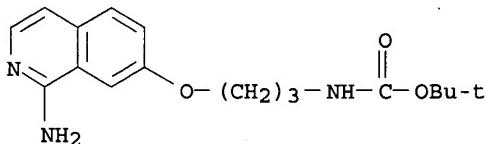
RN 244257-53-2 CAPLUS

CN Carbamic acid, [2-[(1-amino-7-isoquinolinyl)oxy]ethyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



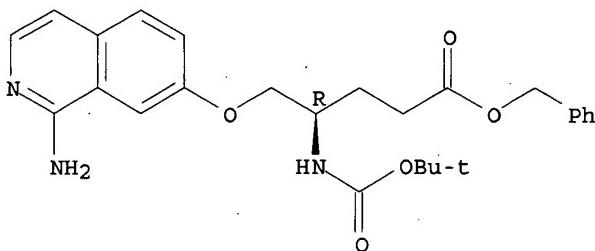
09/ 830,227

RN 244257-58-7 CAPLUS
CN Carbamic acid, [3-[(1-amino-7-isoquinolinyl)oxy]propyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

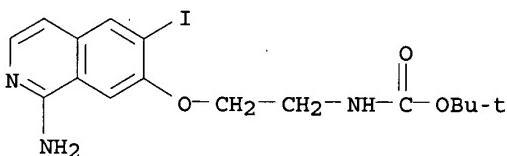


RN 244257-60-1 CAPLUS
CN Pentanoic acid, 5-[(1-amino-7-isoquinolinyl)oxy]-4-[[[(1,1-dimethylethoxy)carbonyl]amino]-, phenylmethyl ester, (4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



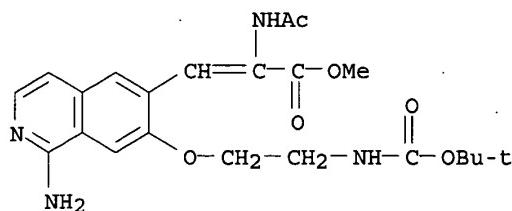
RN 244257-66-7 CAPLUS
CN Carbamic acid, [2-[(1-amino-6-iodo-7-isoquinolinyl)oxy]ethyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



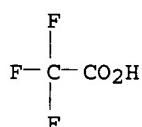
RN 244257-68-9 CAPLUS
CN 2-Propenoic acid, 2-(acetylamino)-3-[1-amino-7-[2-[[[(1,1-dimethylethoxy)carbonyl]amino]ethoxy]-6-isoquinolinyl]-, methyl ester, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-67-8
CMF C22 H28 N4 O6

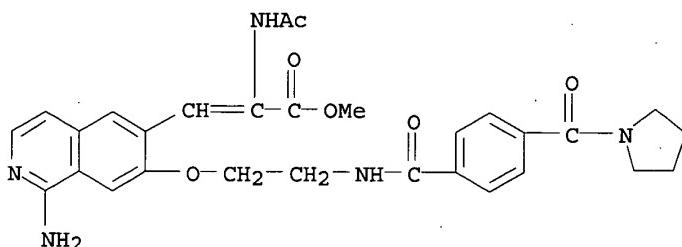


CM 2

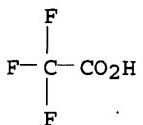
CRN 76-05-1
CMF C2 H F3 O2

RN 244257-70-3 CAPLUS
 CN 2-Propenoic acid, 2-(acetylaminomethyl)-3-[1-amino-7-[[2-[(4-(1-pyrrolidinylcarbonyl)benzoyl)amino]ethoxy]-6-isouinolinyl]-, methyl ester, mono(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 244257-69-0
CMF C29 H31 N5 O6

CM 2

CRN 76-05-1
CMF C2 H F3 O2

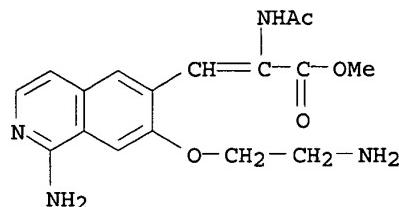
RN 244257-72-5 CAPLUS

09 / 830,227

CN 2-Propenoic acid, 2-(acetylamino)-3-[1-amino-7-(2-aminoethoxy)-6-isooquinolinyl]-, methyl ester, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

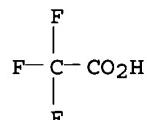
CM 1

CRN 244257-71-4
CMF C17 H20 N4 O4



CM 2

CRN 76-05-1
CMF C2 H F3 O2

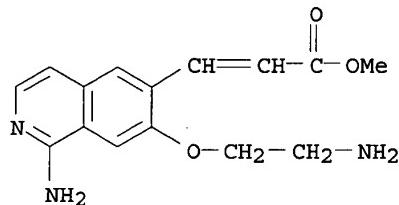


RN 244257-74-7 CAPLUS

CN 2-Propenoic acid, 3-[1-amino-7-(2-aminoethoxy)-6-isoquinolinyl]-, methyl ester, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

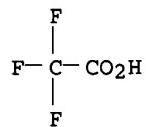
CRN 244257-73-6
CMF C15 H17 N3 O3



CM 2

CRN 76-05-1
CMF C2 H F3 O2

09/ 830,227



REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d his

(FILE 'HOME' ENTERED AT 13:11:32 ON 04 AUG 2003)

FILE 'REGISTRY' ENTERED AT 13:11:41 ON 04 AUG 2003

L1 STRUCTURE uploaded
L2 284 S L1 FUL

FILE 'CAPLUS' ENTERED AT 13:12:06 ON 04 AUG 2003

L3 10 S L2

=> log y

| COST IN U.S. DOLLARS | SINCE FILE ENTRY | TOTAL SESSION |
|----------------------|------------------|---------------|
|----------------------|------------------|---------------|

| | | |
|---------------------|-------|--------|
| FULL ESTIMATED COST | 46.61 | 194.97 |
|---------------------|-------|--------|

| DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) | SINCE FILE ENTRY | TOTAL SESSION |
|--|------------------|---------------|
|--|------------------|---------------|

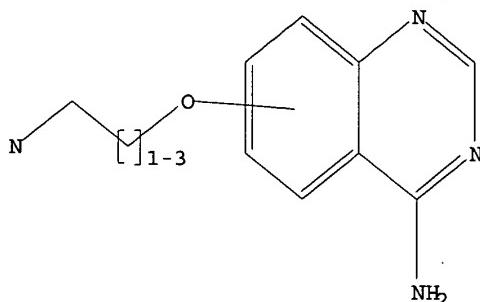
| | | |
|---------------------|-------|-------|
| CA SUBSCRIBER PRICE | -6.51 | -6.51 |
|---------------------|-------|-------|

STN INTERNATIONAL LOGOFF AT 13:14:08 ON 04 AUG 2003

09/ 830,227

L2 STRUCTURE UPLOADED

=> d 12
L2 HAS NO ANSWERS
L2 STR



Structure attributes must be viewed using STN Express query preparation.

=> s 11 ful
FULL SEARCH INITIATED 11:04:44 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 2237 TO ITERATE

100.0% PROCESSED 2237 ITERATIONS 0 ANSWERS
SEARCH TIME: 00.00.03

L3 0 SEA SSS FUL L1

=> s 12 ful
FULL SEARCH INITIATED 11:04:55 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 17635 TO ITERATE

100.0% PROCESSED 17635 ITERATIONS 34 ANSWERS
SEARCH TIME: 00.00.03

L4 34 SEA SSS FUL L2

=> file caplus
COST IN U.S. DOLLARS SINCE FILE TOTAL
FULL ESTIMATED COST ENTRY SESSION
296.30 296.51

FILE 'CAPLUS' ENTERED AT 11:05:04 ON 04 AUG 2003
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

Copyright of the articles to which records in this database refer is held by the publishers listed in the PUBLISHER (PB) field (available for records published or updated in Chemical Abstracts after December 26, 1996), unless otherwise indicated in the original publications. The CA Lexicon is the copyrighted intellectual property of the American Chemical Society and is provided to assist you in searching databases on STN. Any dissemination, distribution, copying, or storing of this information, without the prior written consent of CAS, is strictly prohibited.

FILE COVERS 1907 - 4 Aug 2003 VOL 139 ISS 6

09/ 830,227

FILE LAST UPDATED: 3 Aug 2003 (20030803/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

```
=> s 14  
L5          6 L4
```

```
=> 1- ibib abs hitstr  
1- IS NOT A RECOGNIZED COMMAND  
The previous command name entered was not recognized by the system.  
For a list of commands available to you in the current file, enter  
"HELP COMMANDS" at an arrow prompt (>).
```

```
=> d 14 1- ibib abs hitstr  
YOU HAVE REQUESTED DATA FROM FILE 'REGISTRY' - CONTINUE? (Y)/N:y
```

```
'IBIB' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'  
'ABS' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'  
'HITSTR' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'
```

The following are valid formats:

Substance information can be displayed by requesting individual fields or predefined formats. The predefined substance formats are: (RN = CAS Registry Number)

| | |
|--------|---|
| REG | - RN |
| SAM | - Index Name, MF, and structure - no RN |
| FIDE | - All substance data, except sequence data |
| IDE | - FIDE, but only 50 names |
| SQIDE | - IDE, plus sequence data |
| SQIDE3 | - Same as SQIDE, but 3-letter amino acid codes are used |
| SQD | - Protein sequence data, includes RN |
| SQD3 | - Same as SQD, but 3-letter amino acid codes are used |
| SQN | - Protein sequence name information, includes RN |
| CALC | - Table of calculated properties |
| EPROP | - Table of experimental properties |
| PROP | - EPROP and CALC |

Any CA File format may be combined with any substance format to obtain CA references citing the substance. The substance formats must be cited first. The CA File predefined formats are:

| | |
|-------|--|
| ABS | -- Abstract |
| APPS | -- Application and Priority Information |
| BIB | -- CA Accession Number, plus Bibliographic Data |
| CAN | -- CA Accession Number |
| CBIB | -- CA Accession Number, plus Bibliographic Data (compressed) |
| IND | -- Index Data |
| IPC | -- International Patent Classification |
| PATS | -- PI, SO |
| STD | -- BIB, IPC, and NCL |
| IABS | --ABS, indented, with text labels |
| IBIB | -- BIB, indented, with text labels |
| ISTD | -- STD format, indented |
| OBIB | ----- AN, plus Bibliographic Data (original) |
| OIBIB | ----- OBIB, indented with text labels |

SBIB ----- BIB, no citations
SIBIB ----- IBIB, no citations

The ALL format gives FIDE BIB ABS IND RE, plus sequence data when it is available.

The MAX format is the same as ALL.

The IALL format is the same as ALL with BIB ABS and IND indented, with text labels.

For additional information, please consult the following help messages:

HELP DFIELDS -- To see a complete list of individual display fields.
HELP FORMATS -- To see detailed descriptions of the predefined formats.
ENTER DISPLAY FORMAT (IDE):ibib abs hitstr
'IBIB' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'
'ABS' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'
'HITSTR' IS NOT A VALID FORMAT FOR FILE 'REGISTRY'

The following are valid formats:

Substance information can be displayed by requesting individual fields or predefined formats. The predefined substance formats are: (RN = CAS Registry Number)

REG - RN
SAM - Index Name, MF, and structure - no RN
FIDE - All substance data, except sequence data
IDE - FIDE, but only 50 names
SQIDE - IDE, plus sequence data
SQIDE3 - Same as SQIDE, but 3-letter amino acid codes are used
SQD - Protein sequence data, includes RN
SQD3 - Same as SQD, but 3-letter amino acid codes are used
SQN - Protein sequence name information, includes RN

CALC - Table of calculated properties
EPROP - Table of experimental properties
PROP - EPROP and CALC

Any CA File format may be combined with any substance format to obtain CA references citing the substance. The substance formats must be cited first. The CA File predefined formats are:

ABS -- Abstract
APPS -- Application and Priority Information
BIB -- CA Accession Number, plus Bibliographic Data
CAN -- CA Accession Number
CBIB -- CA Accession Number, plus Bibliographic Data (compressed)
IND -- Index Data
IPC -- International Patent Classification
PATS -- PI, SO
STD -- BIB, IPC, and NCL

IABS -- ABS, indented, with text labels
IBIB -- BIB, indented, with text labels
ISTD -- STD format, indented

OBIB ----- AN, plus Bibliographic Data (original)
OIBIB ----- OBIB, indented with text labels

SBIB ----- BIB, no citations
SIBIB ----- IBIB, no citations

09/ 830,227

The ALL format gives FIDE BIB ABS IND RE, plus sequence data when it is available.

The MAX format is the same as ALL.

The IALL format is the same as ALL with BIB ABS and IND indented, with text labels.

For additional information, please consult the following help messages:

HELP DFIELDS -- To see a complete list of individual display fields.
HELP FORMATS -- To see detailed descriptions of the predefined formats.
ENTER DISPLAY FORMAT (IDE):.
YOU HAVE REQUESTED DATA FROM 34 ANSWERS - CONTINUE? Y/(N):n

=> d his

(FILE 'HOME' ENTERED AT 11:03:53 ON 04 AUG 2003)

FILE 'REGISTRY' ENTERED AT 11:04:04 ON 04 AUG 2003
L1 STRUCTURE uploaded
L2 STRUCTURE uploaded
L3 0 S L1 FUL
L4 34 S L2 FUL

FILE 'CAPLUS' ENTERED AT 11:05:04 ON 04 AUG 2003
L5 6 S L4

FILE 'REGISTRY' ENTERED AT 11:05:30 ON 04 AUG 2003

FILE 'CAPLUS' ENTERED AT 11:05:56 ON 04 AUG 2003

=> d 15 1- ibib abs hitstr
YOU HAVE REQUESTED DATA FROM 6 ANSWERS - CONTINUE? Y/(N):Y

L5 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2003 ACS on STN
ACCESSION NUMBER: 2002:31424 CAPLUS
DOCUMENT NUMBER: 136:102393
TITLE: Preparation of quinazolinylureas for treatment of solid tumors.
PATENT ASSIGNEE(S): AstraZeneca Ab, Swed.; AstraZeneca Uk Ltd.
SOURCE: PCT Int. Appl., 149 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--|----------|-----------------|------------|
| WO 2002002534 | A1 | 20020110 | WO 2001-GB2874 | 20010628 |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |
| AU 2002016758 | A5 | 20020114 | AU 2002-16758 | 20010628 |
| PRIORITY APPLN. INFO.: | | | EP 2000-401897 | A 20000703 |
| | | | WO 2001-GB2874 | W 20010628 |

OTHER SOURCE(S) : MARPAT 136:102393

AB Use of Q1R2NC(:Z)NR3Q2 [Q1 = (substituted) (fused) quinazolinyl, quinolinyl, etc.; Q2 = (substituted) aryl, aralkyl, arylcycloalkyl, heteroaryl, heteroarylalkyl; R2, R3 = H, alkyl; R2R3 = CH₂, CH₂CH₂, (CH₂)₃] as antiinvasive agents in the containment and/or treatment of solid tumor disease is claimed. Thus, 2,6-dichlorophenyl isocyanate was added to a soln. of 4-amino-6-methoxy-7-(N-methylpiperidin-4-ylmethoxy)quinazoline (prepn. given) in CH₂Cl₂/DMF followed by stirring to give 1-(2,6-dichlorophenyl)-3-[6-methoxy-7-(N-methylpiperidin-4-ylmethoxy)quinazolin-4-yl]urea. Title compds. inhibited proliferation of NIH 3T3 fibroblasts with IC₅₀ in the range, for example, of 0.001-10 .mu.M.

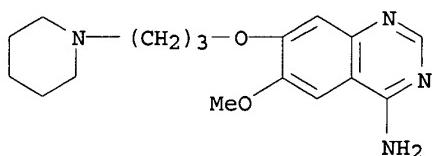
IT 320365-83-1P 320365-84-2P 320365-85-3P
 320365-86-4P 320365-88-6P 320365-89-7P
 320365-91-1P 320365-92-2P 320365-93-3P
 320365-94-4P 320365-95-5P 320365-97-7P
 320366-04-9P 320366-06-1P 320366-08-3P
 320366-10-7P 320366-14-1P 320366-18-5P
 320366-20-9P 320366-24-3P 320366-26-5P
 320366-28-7P 320366-30-1P 320366-31-2P
 320366-64-1P 320366-66-3P 320366-70-9P
 320366-71-0P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(prepn. of quinazolinylureas for treatment of solid tumors)

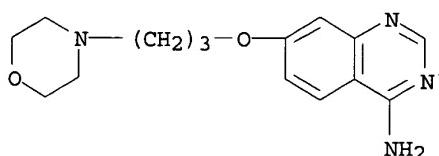
RN 320365-83-1 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(1-piperidinyl)propoxy]- (9CI) (CA INDEX NAME)



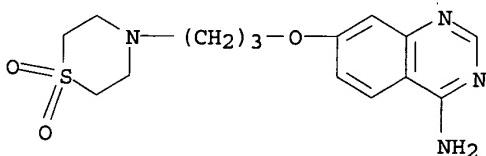
RN 320365-84-2 CAPLUS

CN 4-Quinazolinamine, 7-[3-(4-morpholinyl)propoxy]- (9CI) (CA INDEX NAME)



RN 320365-85-3 CAPLUS

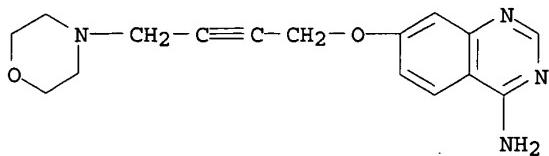
CN 4-Quinazolinamine, 7-[3-(1,1-dioxido-4-thiomorpholinyl)propoxy]- (9CI) (CA INDEX NAME)



09/ 830,227

RN 320365-86-4 CAPLUS

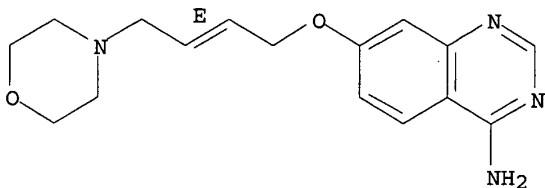
CN 4-Quinazolinamine, 7-[[4-(4-morpholinyl)-2-butynyl]oxy]- (9CI) (CA INDEX NAME)



RN 320365-88-6 CAPLUS

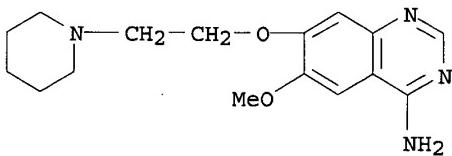
CN 4-Quinazolinamine, 7-[[[(2E)-4-(4-morpholinyl)-2-butenyl]oxy]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.



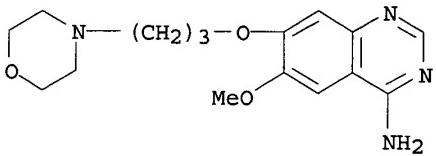
RN 320365-89-7 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[2-(1-piperidinyl)ethoxy]- (9CI) (CA INDEX NAME)



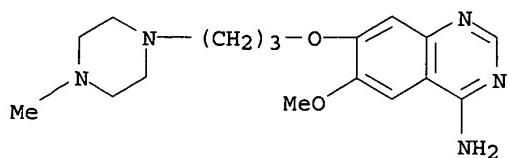
RN 320365-91-1 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(4-morpholinyl)propoxy]- (9CI) (CA INDEX NAME)

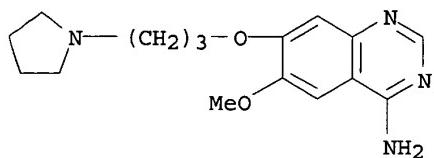


RN 320365-92-2 CAPLUS

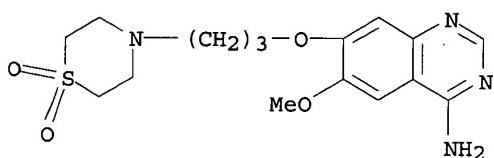
CN 4-Quinazolinamine, 6-methoxy-7-[3-(4-methyl-1-piperazinyl)propoxy]- (9CI) (CA INDEX NAME)



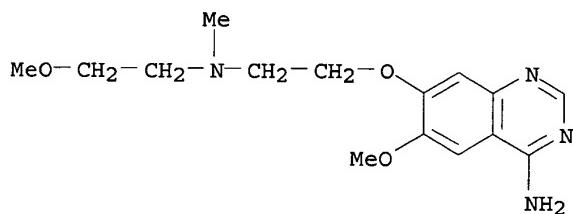
RN 320365-93-3 CAPLUS
CN 4-Quinazolinamine, 6-methoxy-7-[3-(1-pyrrolidinyl)propoxy]- (9CI) (CA INDEX NAME)



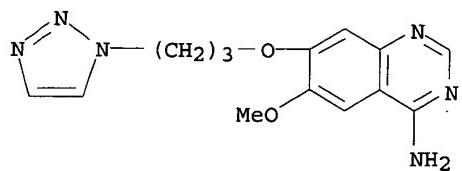
RN 320365-94-4 CAPLUS
CN 4-Quinazolinamine, 7-[3-(1,1-dioxido-4-thiomorpholinyl)propoxy]-6-methoxy- (9CI) (CA INDEX NAME)



RN 320365-95-5 CAPLUS
CN 4-Quinazolinamine, 6-methoxy-7-[2-[(2-methoxyethyl)methylamino]ethoxy]- (9CI) (CA INDEX NAME)

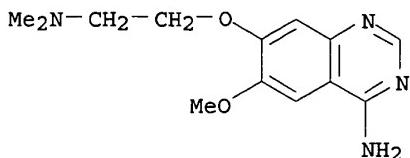


RN 320365-97-7 CAPLUS
CN 4-Quinazolinamine, 6-methoxy-7-[3-(1H-1,2,3-triazol-1-yl)propoxy]- (9CI) (CA INDEX NAME)

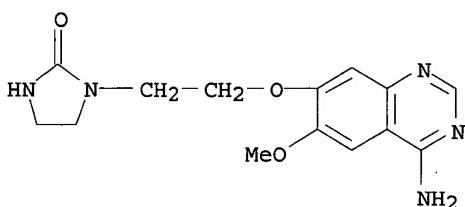


09/ 830,227

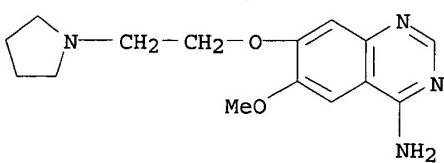
RN 320366-04-9 CAPLUS
CN 4-Quinazolinamine, 7-[2-(dimethylamino)ethoxy]-6-methoxy- (9CI) (CA INDEX NAME)



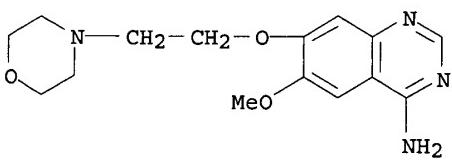
RN 320366-06-1 CAPLUS
CN 2-Imidazolidinone, 1-[2-[(4-amino-6-methoxy-7-quinazolinyl)oxy]ethyl]- (9CI) (CA INDEX NAME)



RN 320366-08-3 CAPLUS
CN 4-Quinazolinamine, 6-methoxy-7-[2-(1-pyrrolidinyl)ethoxy]- (9CI) (CA INDEX NAME)

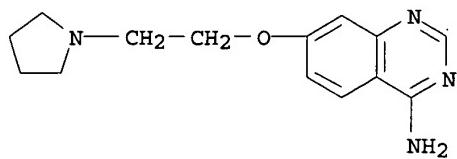


RN 320366-10-7 CAPLUS
CN 4-Quinazolinamine, 6-methoxy-7-[2-(4-morpholinyl)ethoxy]- (9CI) (CA INDEX NAME)

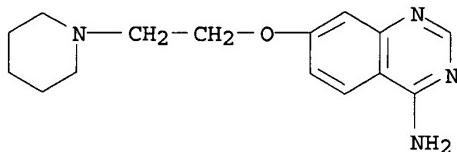


RN 320366-14-1 CAPLUS
CN 4-Quinazolinamine, 7-[2-(1-pyrrolidinyl)ethoxy]- (9CI) (CA INDEX NAME)

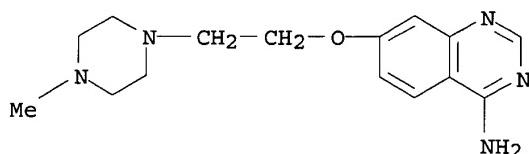
- - - - -



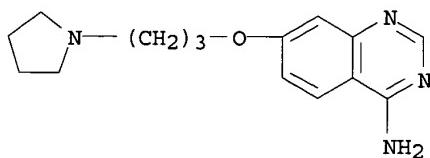
RN 320366-18-5 CAPLUS
CN 4-Quinazolinamine, 7-[2-(1-piperidinyl)ethoxy]- (9CI) (CA INDEX NAME)



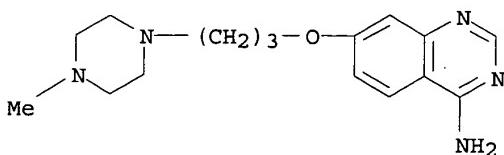
RN 320366-20-9 CAPLUS
CN 4-Quinazolinamine, 7-[2-(4-methyl-1-piperazinyl)ethoxy]- (9CI) (CA INDEX NAME)



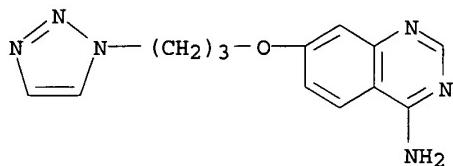
RN 320366-24-3 CAPLUS
CN 4-Quinazolinamine, 7-[3-(1-pyrrolidinyl)propoxy]- (9CI) (CA INDEX NAME)



RN 320366-26-5 CAPLUS
CN 4-Quinazolinamine, 7-[3-(4-methyl-1-piperazinyl)propoxy]- (9CI) (CA INDEX NAME)

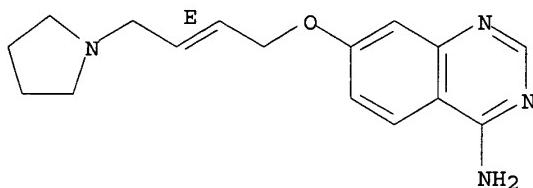


RN 320366-28-7 CAPLUS
CN 4-Quinazolinamine, 7-[3-(1H-1,2,3-triazol-1-yl)propoxy]- (9CI) (CA INDEX NAME)

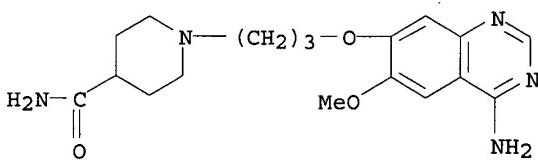


RN 320366-30-1 CAPLUS
CN 4-Quinazolinamine, 7-[[[(2E)-4-(1-pyrrolidinyl)-2-butenyl]oxy]- (9CI) (CA INDEX NAME)

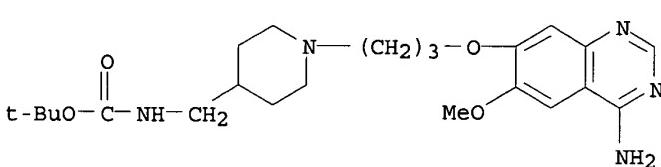
Double bond geometry as shown.



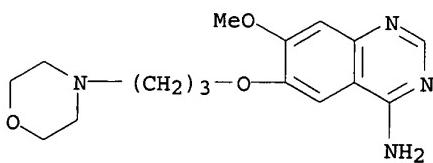
RN 320366-31-2 CAPLUS
CN 4-Piperidinecarboxamide, 1-[3-[(4-amino-6-methoxy-7-quinazolinyl)oxy]propyl]- (9CI) (CA INDEX NAME)



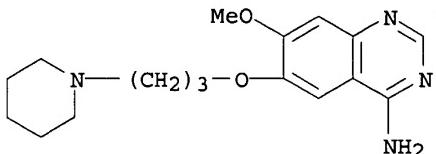
RN 320366-64-1 CAPLUS
CN Carbamic acid, [[1-[3-[(4-amino-6-methoxy-7-quinazolinyl)oxy]propyl]-4-piperidinyl]methyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



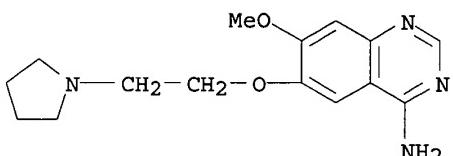
RN 320366-66-3 CAPLUS
CN 4-Quinazolinamine, 7-methoxy-6-[3-(4-morpholinyl)propoxy]- (9CI) (CA INDEX NAME)



RN 320366-70-9 CAPLUS
 CN 4-Quinazolinamine, 7-methoxy-6-[3-(1-piperidinyl)propoxy] - (9CI) (CA
 INDEX NAME)



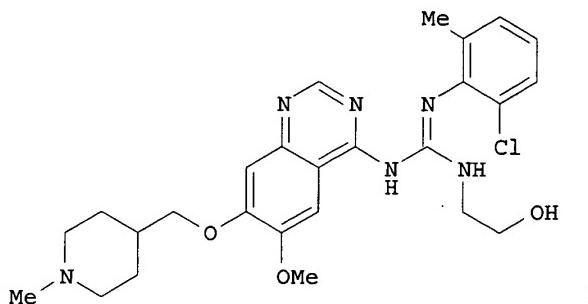
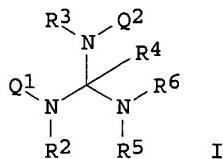
RN 320366-71-0 CAPLUS
 CN 4-Quinazolinamine, 7-methoxy-6-[2-(1-pyrrolidinyl)ethoxy] - (9CI) (CA
 INDEX NAME)



REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 2002:10463 CAPLUS
 DOCUMENT NUMBER: 136:85816
 TITLE: Synthesis of guanidine derivatives of quinazoline and quinoline for use in the treatment of autoimmune diseases
 INVENTOR(S): Poyser, Jeffrey Philip
 PATENT ASSIGNEE(S): AstraZeneca AB, Swed.; AstraZeneca UK Limited
 SOURCE: PCT Int. Appl., 150 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--|----------|-----------------|------------|
| WO 2002000644 | A1 | 20020103 | WO 2001-GB2698 | 20010619 |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |
| EP 1296973 | A1 | 20030402 | EP 2001-940757 | 20010619 |
| R: | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | |
| PRIORITY APPLN. INFO.: | | | GB 2000-15376 | A 20000624 |
| | | | GB 2000-30989 | A 20001219 |
| | | | WO 2001-GB2698 | W 20010619 |



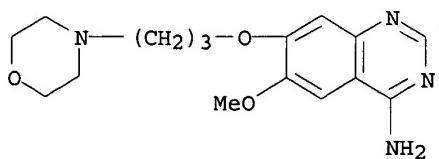
AB Title compds. I [Q1 = (un)substituted quinazolinyl and quinazolinyl-like ring; R2 = H, alkyl; R3 = H, alkyl, or R2 and R3 together form a CH₂, (CH₂)₂ or (CH₂)₃ group; R5 = H, alkyl, or R5 and R6 together with the N atom to which they are attached form a 4- to 7-membered heterocyclic ring optionally contg. a further heteroatom selected from O, N and S, provided that one of the pairs of groups R2 and R4 together, R3 and R4 together and R5 and R4 together forms a bond; Q2 = aryl, arylalkyl, arylcycloalkyl, heteroaryl, heteroarylalkyl or heteroarylcycloalkyl; R6 = (un)substituted group selected from alkenyl, alkynyl, cycloalkyl and cycloalkenyl, or R6 is a substituted alkyl group, and wherein adjacent carbon atoms in any alkylene chain within a R6 group are optionally sepd. by the insertion into the chain of a group selected from O, S, SO, SO₂, amino, CO, etc.; or a tautomer thereof] were prep'd. Over 100 synthetic examples were provided. E.g., Et 3-methoxy-4-((N-methylpiperidin-4-yl)methoxy)benzoate (prepn. given) was nitrated (CH₂Cl₂, TFA, HNO₃, 0.degree.C), the nitro group reduced (MeOH, Pt/C, 1.8 atm H₂), the product condensed/cyclized (2-methoxyethanol, 115.degree.C, 2 h) and treated with thionyl chloride to give 4-chloro-6-methoxy-7-((N-methylpiperidin-4-yl)methoxy)quinazoline. This intermediate was treated with 4-bromo-2-fluorophenol (DMF, K₂CO₃, 100.degree.C, 2.5 h), ammonia in isopropanol (2M, 130.degree.C, 16 h) to give the 4-aminoquinazoline deriv. which was reacted with 2-chloro-6-methylphenylisothiocyanate (DMF, NaH) to afford 1-(2-chloro-6-methylphenyl)-3-[6-methoxy-7-((N-methylpiperidin-4-yl)methoxy)quinazolin-4-yl]thiourea. The thiourea was treated with 2-aminoethanol (CHCl₃/MeOH, HgO, 2 h) to give example compd. II. I are used in the prevention or treatment of T cell mediated diseases.

IT 320365-91-1P, 4-Amino-6-methoxy-7-(3-morpholinopropoxy)quinazoline
320365-93-3P, 4-Amino-6-methoxy-7-(3-(pyrrolidin-1-yl)propoxy)quinazoline 320366-08-3P, 4-Amino-6-methoxy-7-(2-(pyrrolidin-1-yl)ethoxy)quinazoline 320366-10-7P,
4-Amino-6-methoxy-7-(2-morpholinoethoxy)quinazoline 385814-23-3P,
, 4-Amino-6-methoxy-7-(2-pyridylmethoxy)quinazoline 385814-97-1P

, 4-Amino-7-(2-morpholinoethoxy)quinazoline
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
(Reactant or reagent)
(intermediate; synthesis of guanidine derivs. of quinazoline and
quinoline for use in treatment of autoimmune diseases)

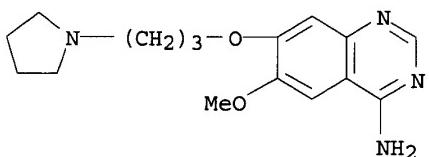
RN 320365-91-1 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(4-morpholinyl)propoxy]- (9CI) (CA
INDEX NAME)



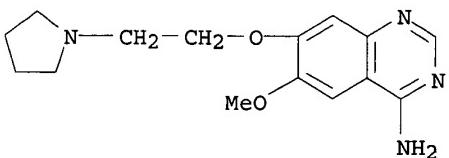
RN 320365-93-3 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(1-pyrrolidinyl)propoxy]- (9CI) (CA
INDEX NAME)



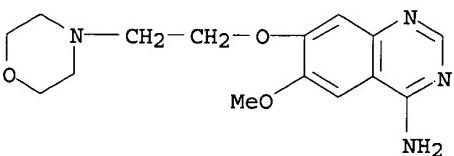
RN 320366-08-3 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[2-(1-pyrrolidinyl)ethoxy]- (9CI) (CA
INDEX NAME)



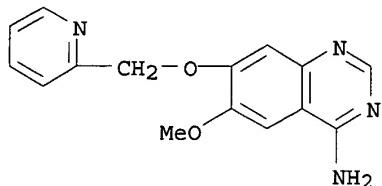
RN 320366-10-7 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[2-(4-morpholinyl)ethoxy]- (9CI) (CA INDEX
NAME)

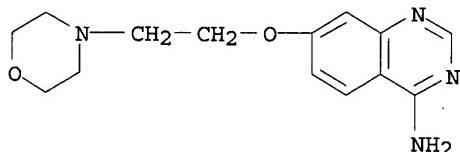


RN 385814-23-3 CAPLUS

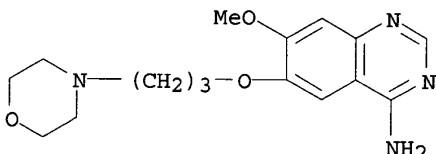
CN 4-Quinazolinamine, 6-methoxy-7-(2-pyridinylmethoxy)- (9CI) (CA INDEX
NAME)



RN 385814-97-1 CAPLUS
 CN 4-Quinazolinamine, 7-[2-(4-morpholinyl)ethoxy]- (9CI) (CA INDEX NAME)



IT 320366-66-3, 4-Amino-7-methoxy-6-(3-morpholinopropoxy)quinazoline
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (reactant; synthesis of guanidine derivs. of quinazoline and quinoline
 for use in treatment of autoimmune diseases)
 RN 320366-66-3 CAPLUS
 CN 4-Quinazolinamine, 7-methoxy-6-[3-(4-morpholinyl)propoxy]- (9CI) (CA
 INDEX NAME)



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 2001:676589 CAPLUS
 DOCUMENT NUMBER: 135:227013
 TITLE: Preparation of quinazolinylureas and analogs as VEGF
 receptor antagonists
 INVENTOR(S): Hennequin, Laurent Francois Andre; Crawley, Graham
 Charles; McKerrecher, Darren; Ple, Patrick; Poyser,
 Jeffrey Philip; Lambert, Christine Marie Paul
 PATENT ASSIGNEE(S): AstraZeneca AB, Swed.; AstraZeneca UK Limited
 SOURCE: PCT Int. Appl., 170 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---------------|------|----------|-----------------|----------|
| WO 2001066099 | A2 | 20010913 | WO 2001-GB863 | 20010301 |
| WO 2001066099 | A3 | 20020321 | | |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,

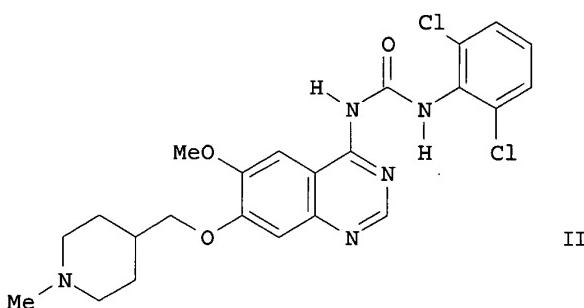
CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
 HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
 LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
 SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
 YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
 BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG

EP 1272185 A2 20030108 EP 2001-907938 20010301

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR

PRIORITY APPLN. INFO.: EP 2000-400595 A 20000306
 WO 2001-GB863 W 20010301

OTHER SOURCE(S) : MARPAT 135:227013
 GI



AB Q1NR2C(:X)NR3Q2 [I; Q1 = e.g., (un)substituted 4-quinazolinyl; Q2 = (un)substituted (hetero)aryl(alkyl), cycloalkyl, etc.; R2,R3 = H or alkyl; R2R3 = (CH₂)₁₋₃; X = O, S; NCN, (alkyl)imino] were prep'd. Thus, Et piperidine-4-carboxylate was converted in 7 steps to Et 2-amino-5-methoxy-4-(1-methylpiperidine-4-ylmethoxy)benzoate which was cyclocondensed with HC(:NH)NH₂.HOAc and the product converted in 4 steps to title compd. II. Data for biol. activity of I were given.

IT 320365-83-1P, 4-Amino-6-methoxy-7-(3-piperidinopropoxy)quinazoline

320365-84-2P 320365-85-3P, 4-Amino-7-[3-(1,1-dioxotetrahydro-4H-1,4-thiazin-4-yl)propoxy]quinazoline

320365-86-4P 320365-88-6P 320365-89-7P

320365-91-1P 320365-92-2P 320365-93-3P

320365-94-4P 320365-95-5P 320365-97-7P

320366-04-9P 320366-06-1P 320366-08-3P

320366-10-7P 320366-14-1P 320366-18-5P

320366-20-9P 320366-24-3P 320366-26-5P

320366-28-7P 320366-30-1P 320366-31-2P

320366-64-1P 320366-66-3P 320366-70-9P

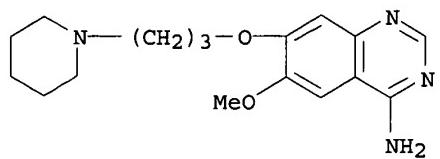
320366-71-0P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

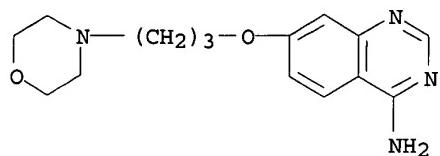
(prepn. of quinazolinylureas and analogs as VEGF receptor antagonists)

RN 320365-83-1 CAPLUS

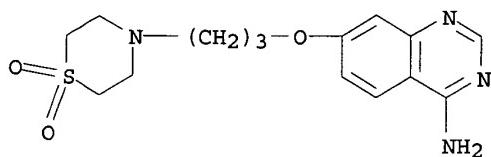
CN 4-Quinazolinamine, 6-methoxy-7-[3-(1-piperidinyl)propoxy]- (9CI) (CA INDEX NAME)



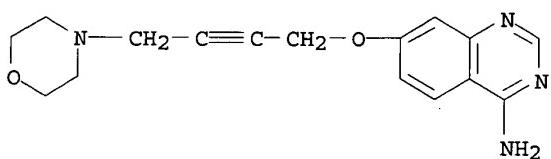
RN 320365-84-2 CAPLUS
CN 4-Quinazolinamine, 7-[3-(4-morpholinyl)propoxy]- (9CI) (CA INDEX NAME)



RN 320365-85-3 CAPLUS
CN 4-Quinazolinamine, 7-[3-(1,1-dioxido-4-thiomorpholinyl)propoxy]- (9CI)
(CA INDEX NAME)

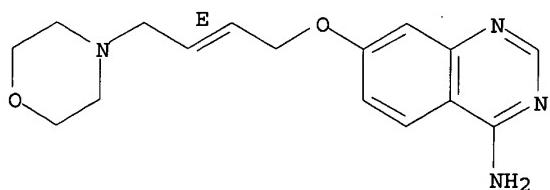


RN 320365-86-4 CAPLUS
CN 4-Quinazolinamine, 7-[4-(4-morpholinyl)-2-butynyl]oxy]- (9CI) (CA INDEX NAME)



RN 320365-88-6 CAPLUS
CN 4-Quinazolinamine, 7-[[2E)-4-(4-morpholinyl)-2-butenyl]oxy]- (9CI) (CA INDEX NAME)

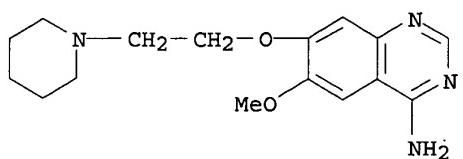
Double bond geometry as shown.



RN 320365-89-7 CAPLUS

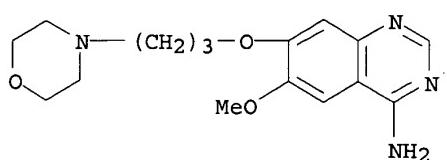
09/ 830,227

CN 4-Quinazolinamine, 6-methoxy-7-[2-(1-piperidinyl)ethoxy]- (9CI) (CA INDEX NAME)



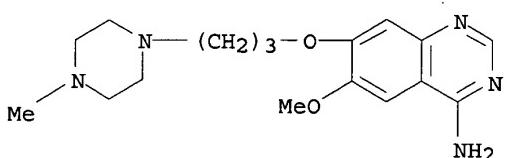
RN 320365-91-1 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(4-morpholinyl)propoxy]- (9CI) (CA INDEX NAME)



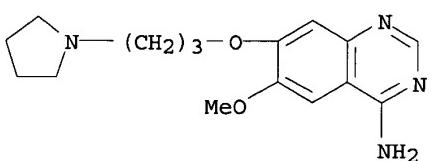
RN 320365-92-2 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(4-methyl-1-piperazinyl)propoxy]- (9CI) (CA INDEX NAME)



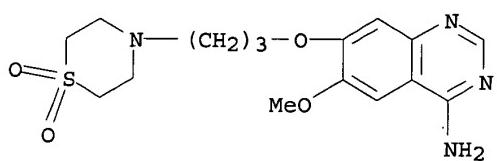
RN 320365-93-3 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(1-pyrrolidinyl)propoxy]- (9CI) (CA INDEX NAME)



RN 320365-94-4 CAPLUS

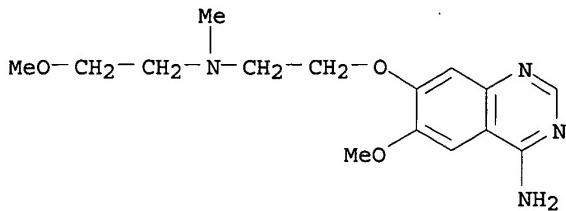
CN 4-Quinazolinamine, 7-[3-(1,1-dioxido-4-thiomorpholinyl)propoxy]-6-methoxy- (9CI) (CA INDEX NAME)



09/ 830,227

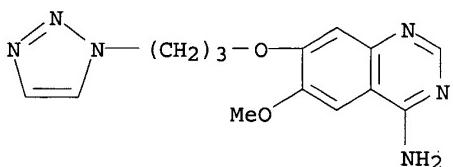
RN 320365-95-5 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[2-[(2-methoxyethyl)methylamino]ethoxy]-
(9CI) (CA INDEX NAME)



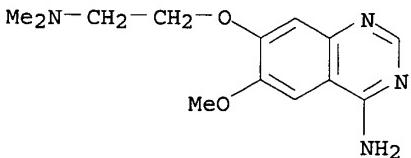
RN 320365-97-7 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(1H-1,2,3-triazol-1-yl)propoxy]- (9CI)
(CA INDEX NAME)



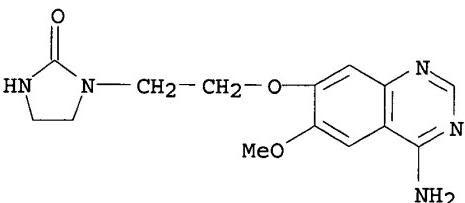
RN 320366-04-9 CAPLUS

CN 4-Quinazolinamine, 7-[2-(dimethylamino)ethoxy]-6-methoxy- (9CI) (CA INDEX
NAME)



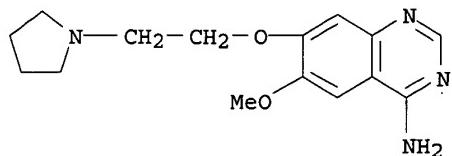
RN 320366-06-1 CAPLUS

CN 2-Imidazolidinone, 1-[2-[(4-amino-6-methoxy-7-quinazolinyl)oxy]ethyl]-
(9CI) (CA INDEX NAME)

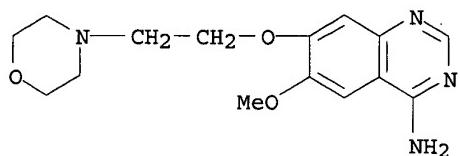


RN 320366-08-3 CAPLUS

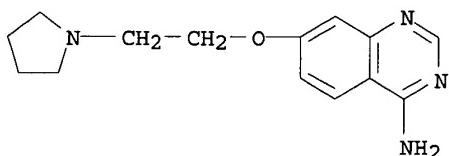
CN 4-Quinazolinamine, 6-methoxy-7-[2-(1-pyrrolidinyl)ethoxy]- (9CI) (CA
INDEX NAME)



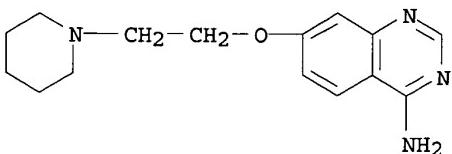
RN 320366-10-7 CAPLUS
CN 4-Quinazolinamine, 6-methoxy-7-[2-(4-morpholinyl)ethoxy]- (9CI) (CA INDEX NAME)



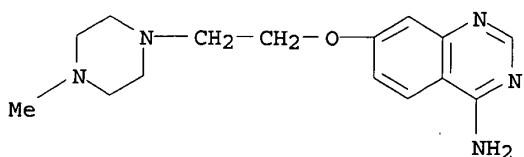
RN 320366-14-1 CAPLUS
CN 4-Quinazolinamine, 7-[2-(1-pyrrolidinyl)ethoxy]- (9CI) (CA INDEX NAME)



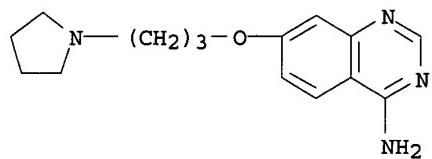
RN 320366-18-5 CAPLUS
CN 4-Quinazolinamine, 7-[2-(1-piperidinyl)ethoxy]- (9CI) (CA INDEX NAME)



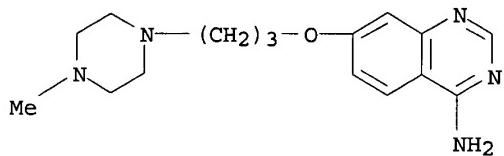
RN 320366-20-9 CAPLUS
CN 4-Quinazolinamine, 7-[2-(4-methyl-1-piperazinyl)ethoxy]- (9CI) (CA INDEX NAME)



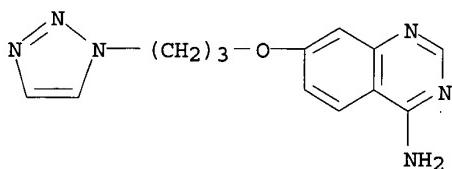
RN 320366-24-3 CAPLUS
CN 4-Quinazolinamine, 7-[3-(1-pyrrolidinyl)propoxy]- (9CI) (CA INDEX NAME)



RN 320366-26-5 CAPLUS
CN 4-Quinazolinamine, 7-[3-(4-methyl-1-piperazinyl)propoxy]- (9CI) (CA INDEX NAME)

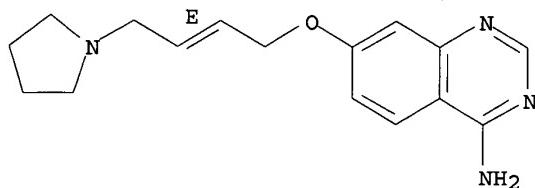


RN 320366-28-7 CAPLUS
CN 4-Quinazolinamine, 7-[3-(1H-1,2,3-triazol-1-yl)propoxy]- (9CI) (CA INDEX NAME)

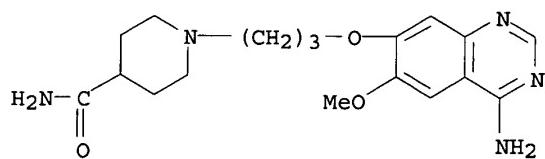


RN 320366-30-1 CAPLUS
CN 4-Quinazolinamine, 7-[[(2E)-4-(1-pyrrolidinyl)-2-butenyl]oxy]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.



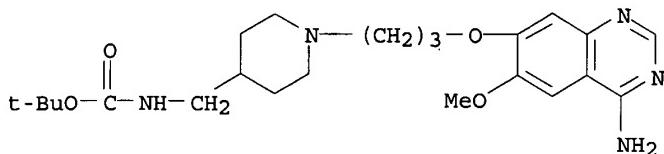
RN 320366-31-2 CAPLUS
CN 4-Piperidinecarboxamide, 1-[3-[(4-amino-6-methoxy-7-quinazolinyl)oxy]propyl]- (9CI) (CA INDEX NAME)



09/ 830,227

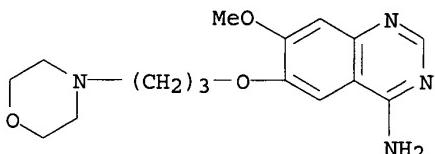
RN 320366-64-1 CAPLUS

CN Carbamic acid, [1-[3-[(4-amino-6-methoxy-7-quinazolinyl)oxy]propyl]-4-piperidinyl]methyl-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



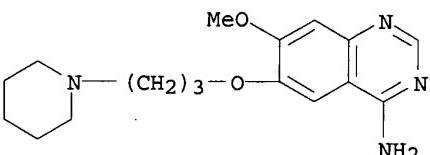
RN 320366-66-3 CAPLUS

CN 4-Quinazolinamine, 7-methoxy-6-[3-(4-morpholinyl)propoxy]- (9CI) (CA INDEX NAME)



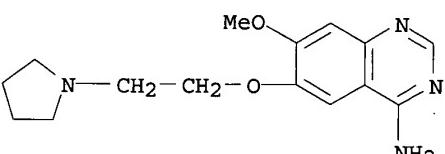
RN 320366-70-9 CAPLUS

CN 4-Quinazolinamine, 7-methoxy-6-[3-(1-piperidinyl)propoxy]- (9CI) (CA INDEX NAME)



RN 320366-71-0 CAPLUS

CN 4-Quinazolinamine, 7-methoxy-6-[2-(1-pyrrolidinyl)ethoxy]- (9CI) (CA INDEX NAME)



L5 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 2001:50631 CAPLUS

DOCUMENT NUMBER: 134:100885

TITLE: Preparation of quinazolinyl ureas, thioureas and guanidines for use in the prevention or treatment of T cell mediated diseases or medical conditions

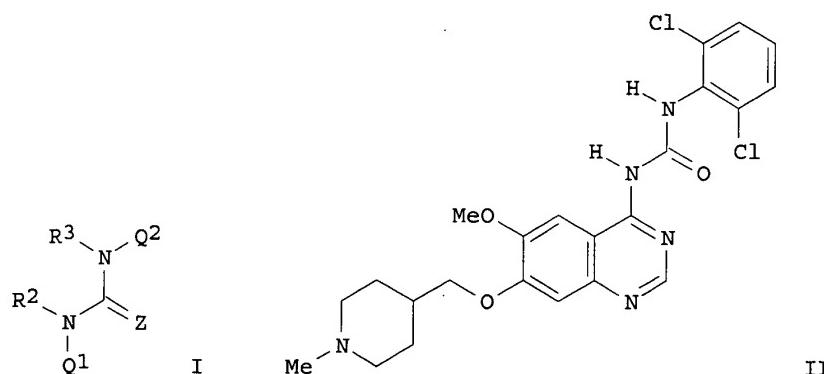
INVENTOR(S): Crawley, Graham Charles; McKerrecher, Darren; Poyser, Jeffrey Philip; Hennequin, Laurent Francois Andre; Ple, Patrick; Lambert, Christine Marie-Paul

PATENT ASSIGNEE(S) : Astrazeneca UK Limited, UK; Zeneca Pharma S.A.
SOURCE : PCT Int. Appl., 169 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE : English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--|----------|-----------------|------------|
| WO 2001004102 | A1 | 20010118 | WO 2000-GB2566 | 20000704 |
| W: | AE, AG, AL, AM, AT, AU, AZ, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |
| BR 2000012157 | A | 20020402 | BR 2000-12157 | 20000704 |
| EP 1218353 | A1 | 20020703 | EP 2000-953271 | 20000704 |
| R: | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL | | | |
| JP 2003504360 | T2 | 20030204 | JP 2001-509712 | 20000704 |
| NO 2002000042 | A | 20020304 | NO 2002-42 | 20020104 |
| PRIORITY APPLN. INFO.: | | | EP 1999-401692 | A 19990707 |
| | | | EP 2000-401221 | A 20000504 |
| | | | WO 2000-GB2566 | W 20000704 |

OTHER SOURCE(S) : MARPAT 134:100885
GI



AB The title compds. [I; Q1 = quinazoline ring optionally substituted with halo, CF₃ or CN, or a group X₁Q₃ (wherein X₁ = a direct bond, O; Q₃ = aryl, arylalkyl, heterocyclyl, (heterocyclyl)alkyl); R₂, R₃ = H, alkyl; Z = O, S, NH; Q₂ = aryl, arylalkyl] and their pharmaceutically-acceptable salts, useful in the prevention or treatment of T cell mediated diseases or medical conditions such as transplant rejection or rheumatoid arthritis, were prep'd. and formulated. E.g., a multi-step synthesis of the urea II was given. In general, activity possessed by compds. I may be demonstrated at IC₅₀ of 0.0001- 5 .mu.M against enzyme p56lck binding and IC₅₀ of 0.001-10 .mu.M in in vitro T cell proliferation assay (T cell receptor stimulation).

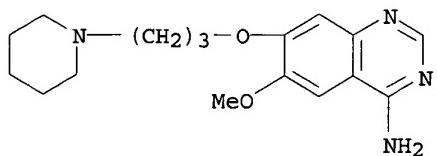
IT 320365-83-1P 320365-84-2P 320365-85-3P
 320365-86-4P 320365-88-6P 320365-89-7P
 320365-91-1P 320365-92-2P 320365-93-3P
 320365-94-4P 320365-95-5P 320365-97-7P
 320366-04-9P 320366-06-1P 320366-08-3P
 320366-10-7P 320366-14-1P 320366-18-5P
 320366-20-9P 320366-24-3P 320366-26-5P
 320366-28-7P 320366-30-1P 320366-31-2P
 320366-64-1P 320366-66-3P 320366-70-9P
 320366-71-0P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(prepn. of quinazolinyl ureas, thioureas and guanidines for use in the prevention or treatment of T cell mediated diseases or medical conditions)

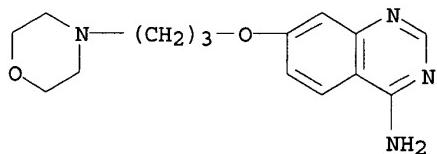
RN 320365-83-1 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(1-piperidinyl)propoxy] - (9CI) (CA INDEX NAME)



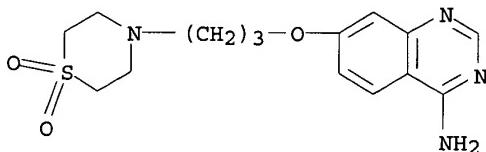
RN 320365-84-2 CAPLUS

CN 4-Quinazolinamine, 7-[3-(4-morpholinyl)propoxy] - (9CI) (CA INDEX NAME)



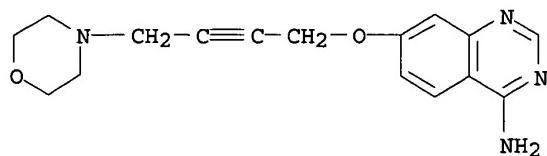
RN 320365-85-3 CAPLUS

CN 4-Quinazolinamine, 7-[3-(1,1-dioxido-4-thiomorpholinyl)propoxy] - (9CI) (CA INDEX NAME)



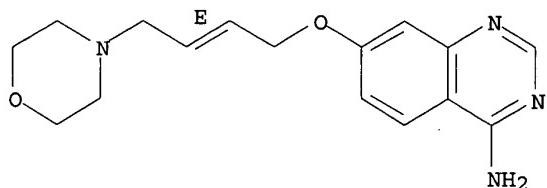
RN 320365-86-4 CAPLUS

CN 4-Quinazolinamine, 7-[[4-(4-morpholinyl)-2-butynyl]oxy] - (9CI) (CA INDEX NAME)

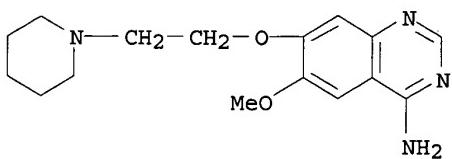


RN 320365-88-6 CAPLUS
CN 4-Quinazolinamine, 7-[(2E)-4-(4-morpholinyl)-2-butenyl]oxy]- (9CI) (CA INDEX NAME)

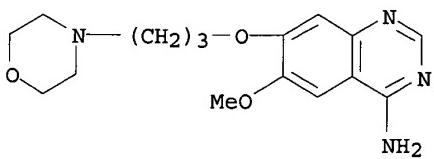
Double bond geometry as shown.



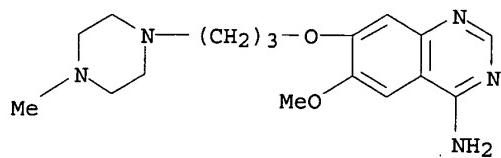
RN 320365-89-7 CAPLUS
CN 4-Quinazolinamine, 6-methoxy-7-[2-(1-piperidinyl)ethoxy]- (9CI) (CA INDEX NAME)



RN 320365-91-1 CAPLUS
CN 4-Quinazolinamine, 6-methoxy-7-[3-(4-morpholinyl)propoxy]- (9CI) (CA INDEX NAME)



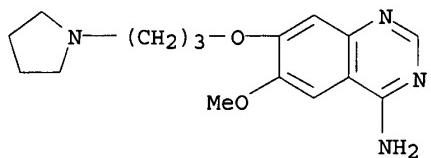
RN 320365-92-2 CAPLUS
CN 4-Quinazolinamine, 6-methoxy-7-[3-(4-methyl-1-piperazinyl)propoxy]- (9CI) (CA INDEX NAME)



09/ 830,227

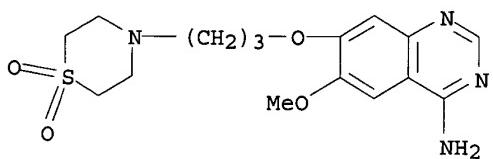
RN 320365-93-3 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(1-pyrrolidinyl)propoxy] - (9CI) (CA INDEX NAME)



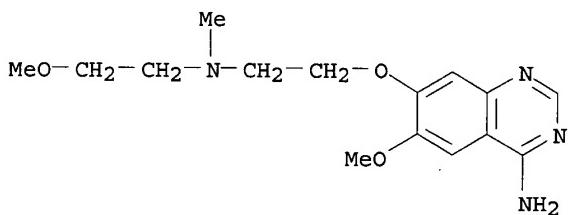
RN 320365-94-4 CAPLUS

CN 4-Quinazolinamine, 7-[3-(1,1-dioxido-4-thiomorpholinyl)propoxy]-6-methoxy- (9CI) (CA INDEX NAME)



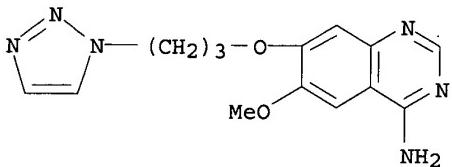
RN 320365-95-5 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[2-[(2-methoxyethyl)methylamino]ethoxy] - (9CI) (CA INDEX NAME)



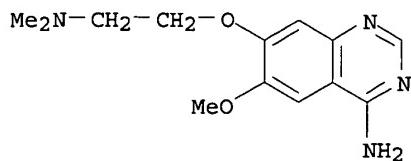
RN 320365-97-7 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[3-(1H-1,2,3-triazol-1-yl)propoxy] - (9CI) (CA INDEX NAME)



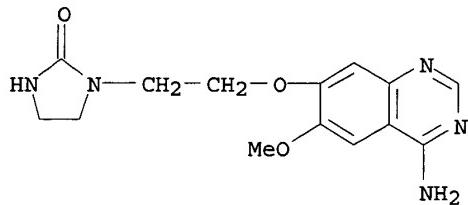
RN 320366-04-9 CAPLUS

CN 4-Quinazolinamine, 7-[2-(dimethylamino)ethoxy]-6-methoxy- (9CI) (CA INDEX NAME)



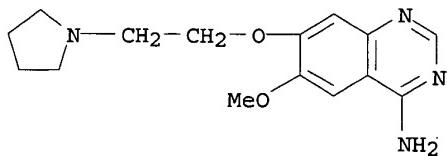
RN 320366-06-1 CAPLUS

CN 2-Imidazolidinone, 1-[2-[(4-amino-6-methoxy-7-quinazolinyl)oxy]ethyl] - (9CI) (CA INDEX NAME)



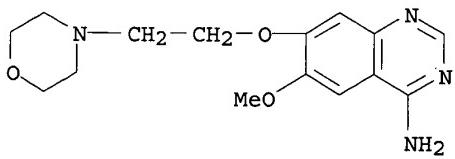
RN 320366-08-3 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[2-(1-pyrrolidinyl)ethoxy] - (9CI) (CA INDEX NAME)



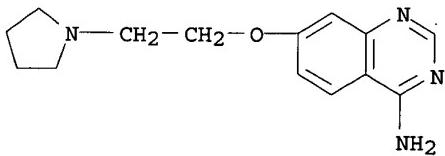
RN 320366-10-7 CAPLUS

CN 4-Quinazolinamine, 6-methoxy-7-[2-(4-morpholinyl)ethoxy] - (9CI) (CA INDEX NAME)



RN 320366-14-1 CAPLUS

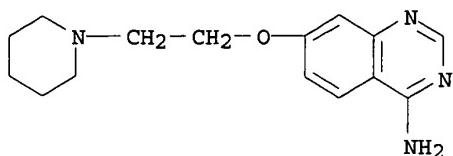
CN 4-Quinazolinamine, 7-[2-(1-pyrrolidinyl)ethoxy] - (9CI) (CA INDEX NAME)



RN 320366-18-5 CAPLUS

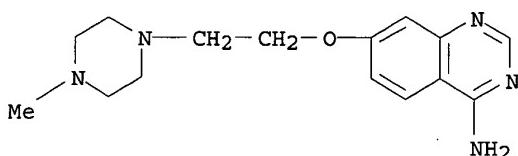
09/ 830,227

CN 4-Quinazolinamine, 7-[2-(1-piperidinyl)ethoxy] - (9CI) (CA INDEX NAME)



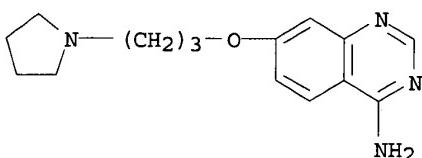
RN 320366-20-9 CAPLUS

CN 4-Quinazolinamine, 7-[2-(4-methyl-1-piperazinyl)ethoxy] - (9CI) (CA INDEX NAME)



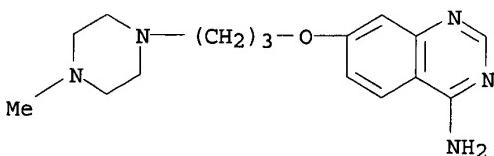
RN 320366-24-3 CAPLUS

CN 4-Quinazolinamine, 7-[3-(1-pyrrolidinyl)propoxy] - (9CI) (CA INDEX NAME)



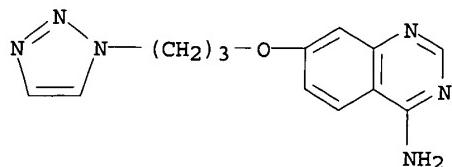
RN 320366-26-5 CAPLUS

CN 4-Quinazolinamine, 7-[3-(4-methyl-1-piperazinyl)propoxy] - (9CI) (CA INDEX NAME)



RN 320366-28-7 CAPLUS

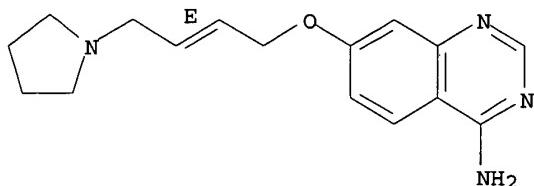
CN 4-Quinazolinamine, 7-[3-(1H-1,2,3-triazol-1-yl)propoxy] - (9CI) (CA INDEX NAME)



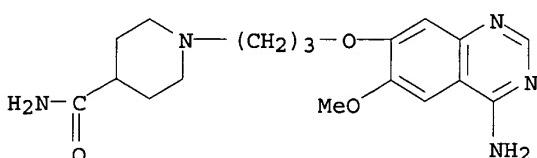
09/ 830,227

RN 320366-30-1 CAPLUS
CN 4-Quinazolinamine, 7-[[(2E)-4-(1-pyrrolidinyl)-2-but enyl]oxy]- (9CI) (CA INDEX NAME)

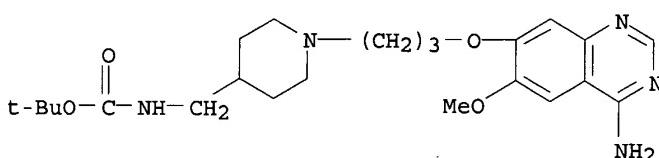
Double bond geometry as shown.



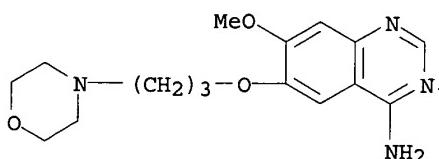
RN 320366-31-2 CAPLUS
CN 4-Piperidinecarboxamide, 1-[3-[(4-amino-6-methoxy-7-quinazolinyl)oxy]propyl]- (9CI) (CA INDEX NAME)



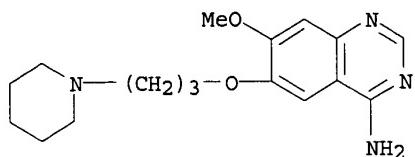
RN 320366-64-1 CAPLUS
CN Carbamic acid, [[1-[3-[(4-amino-6-methoxy-7-quinazolinyl)oxy]propyl]-4-piperidinyl]methyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



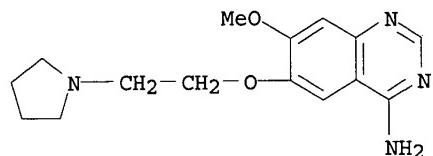
RN 320366-66-3 CAPLUS
CN 4-Quinazolinamine, 7-methoxy-6-[3-(4-morpholinyl)propoxy]- (9CI) (CA INDEX NAME)



RN 320366-70-9 CAPLUS
CN 4-Quinazolinamine, 7-methoxy-6-[3-(1-piperidinyl)propoxy]- (9CI) (CA INDEX NAME)



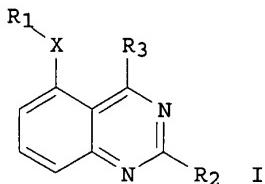
RN 320366-71-0 CAPLUS
 CN 4-Quinazolinamine, 7-methoxy-6-[2-(1-pyrrolidinyl)ethoxy]- (9CI) (CA
 INDEX NAME)



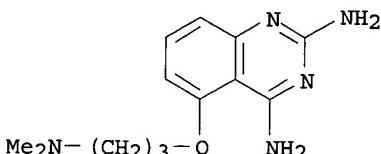
REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 1998:745041 CAPLUS
 DOCUMENT NUMBER: 130:10618
 TITLE: Modulating serine/threonine protein kinase function with quinazoline-based compounds and their use as antitumor and anti-fibrotic agents
 INVENTOR(S): Tang, Peng C.; McMahon, Gerald; Weinberger, Heinz; Kutscher, Bernhard; App, Harald
 PATENT ASSIGNEE(S): Sugen, Inc., USA
 SOURCE: PCT Int. Appl., 147 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|-------------|
| WO 9850370 | A1 | 19981112 | WO 1998-US9060 | 19980501 |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG | | | | |
| ZA 9803669 | A | 19991101 | ZA 1998-3669 | 19980430 |
| AU 9872829 | A1 | 19981127 | AU 1998-72829 | 19980501 |
| EP 981519 | A1 | 20000301 | EP 1998-920203 | 19980501 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI | | | | |
| US 6204267 | B1 | 20010320 | US 1998-71682 | 19980501 |
| JP 2001524128 | T2 | 20011127 | JP 1998-548336 | 19980501 |
| US 2001014679 | A1 | 20010816 | US 2001-769360 | 20010126 |
| PRIORITY APPLN. INFO.: | | | US 1997-45351P | P 19970502 |
| | | | US 1997-60152P | P 19970926 |
| | | | US 1998-71682 | A3 19980501 |



- AB The present invention is directed in part towards methods of modulating the function of serine/threonine protein kinases with quinazoline-based compds (I). The methods incorporate cells that express a serine/threonine protein kinase, such as RAF. In addn., the invention describes methods of preventing and treating serine/threonine protein kinase-related abnormal conditions (e.g., tumors, fibrotic disorders, or other signal transduction aberrations) in organisms with a compd. identified by the invention. Furthermore, the invention pertains to quinazoline compds. and pharmaceutical compns. comprising these compds. Syntheses and biol. activities are provided for 38 quinazoline-based compds.
- IT 215925-99-8P
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (modulating serine/threonine protein kinase function with quinazoline-based compds. and their use as antitumor and anti-fibrotic agents)
- RN 215925-99-8 CAPLUS
- CN 2,4-Quinazoliniediamine, 5-[3-(dimethylamino)propoxy] - (9CI) (CA INDEX NAME)



REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L5 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 1994:315146 CAPLUS
 DOCUMENT NUMBER: 120:315146
 TITLE: The aminoquinazoline group as a replacement for the salicylamide group: The design and synthesis of a novel highly selective .beta.1 adrenoceptor partial agonist
 AUTHOR(S): Block, Michael H.; Kenny, Peter W.; Thomson, David S.; Yu, Man Tat
 CORPORATE SOURCE: ICI Pharm., Alderley Park/Macclesfield/Cheshire, SK10 4TG, UK
 SOURCE: Drug Design and Discovery (1992), 9(2), 167-76, (plate)
 CODEN: DDDIEV; ISSN: 1055-9612

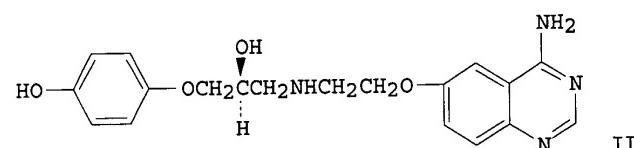
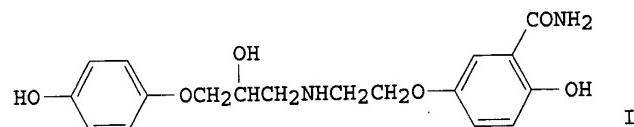
DOCUMENT TYPE:

Journal

LANGUAGE:

English

GI



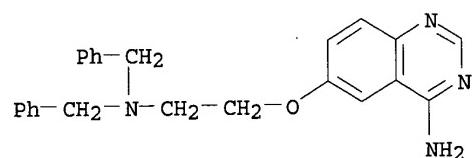
AB The high potency at β .1 receptors, excellent selectivity (β .1/ β .2) and high degree of agonism displayed by compds. such as I is believed to be due in part to the salicylamide side chain. Two conformations of salicylamide are known to exist in the crystal state, but ab initio calcns. suggest that in the absence of crystal packing forces one of them contg. the amide group should be more stable. The aminoquinazoline group was judged to be a good replacement for salicylamide in I, and consequently the oxypropanolamine deriv. (II) was prep'd. II shows extremely high potency at the β .1 receptor, and excellent β .1/ β .2 selectivity. It has comparable in vitro activity to I, although it displays a lower degree of agonism. In this system, aminoquinazoline appears to be an excellent mimic of the salicylamide group.

IT **154664-43-4P**

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(prepn. and hydrogenolysis of)

RN 154664-43-4 CAPLUS

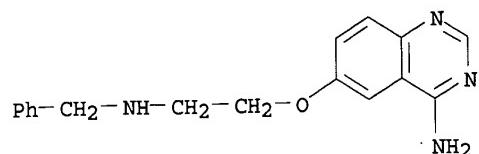
CN 4-Quinazolinamine, 6-[2-[bis(phenylmethyl)amino]ethoxy]- (9CI) (CA INDEX NAME)

IT **154664-42-3P**

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(prepn. and reaction with epoxide deriv.)

RN 154664-42-3 CAPLUS

CN 4-Quinazolinamine, 6-[2-[(phenylmethyl)amino]ethoxy]- (9CI) (CA INDEX NAME)



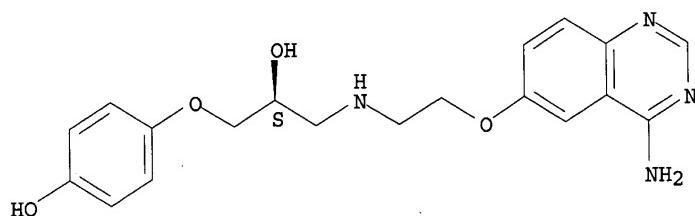
IT 154664-41-2P

RL: SPM (Synthetic preparation); PREP (Preparation)
 (prepn. and .beta.1-adrenergic agonists activity of, structure in
 relation to)

RN 154664-41-2 CAPLUS

CN Phenol, 4-[[3-[(2-[(4-amino-6-quinazolinyl)oxy]ethyl)amino]-2-hydroxypropoxy]-, (S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



=> d his

(FILE 'HOME' ENTERED AT 11:03:53 ON 04 AUG 2003)

FILE 'REGISTRY' ENTERED AT 11:04:04 ON 04 AUG 2003

L1 STRUCTURE uploaded
 L2 STRUCTURE uploaded
 L3 0 S L1 FUL
 L4 34 S L2 FUL

FILE 'CAPLUS' ENTERED AT 11:05:04 ON 04 AUG 2003

L5 6 S L4

FILE 'REGISTRY' ENTERED AT 11:05:30 ON 04 AUG 2003

FILE 'CAPLUS' ENTERED AT 11:05:56 ON 04 AUG 2003

=> log y

| COST IN U.S. DOLLARS | SINCE FILE ENTRY | TOTAL SESSION |
|--|------------------|---------------|
| FULL ESTIMATED COST | 27.63 | 324.96 |
| DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) | SINCE FILE ENTRY | TOTAL SESSION |
| CA SUBSCRIBER PRICE | -3.91 | -3.91 |

STN INTERNATIONAL LOGOFF AT 11:06:47 ON 04 AUG 2003